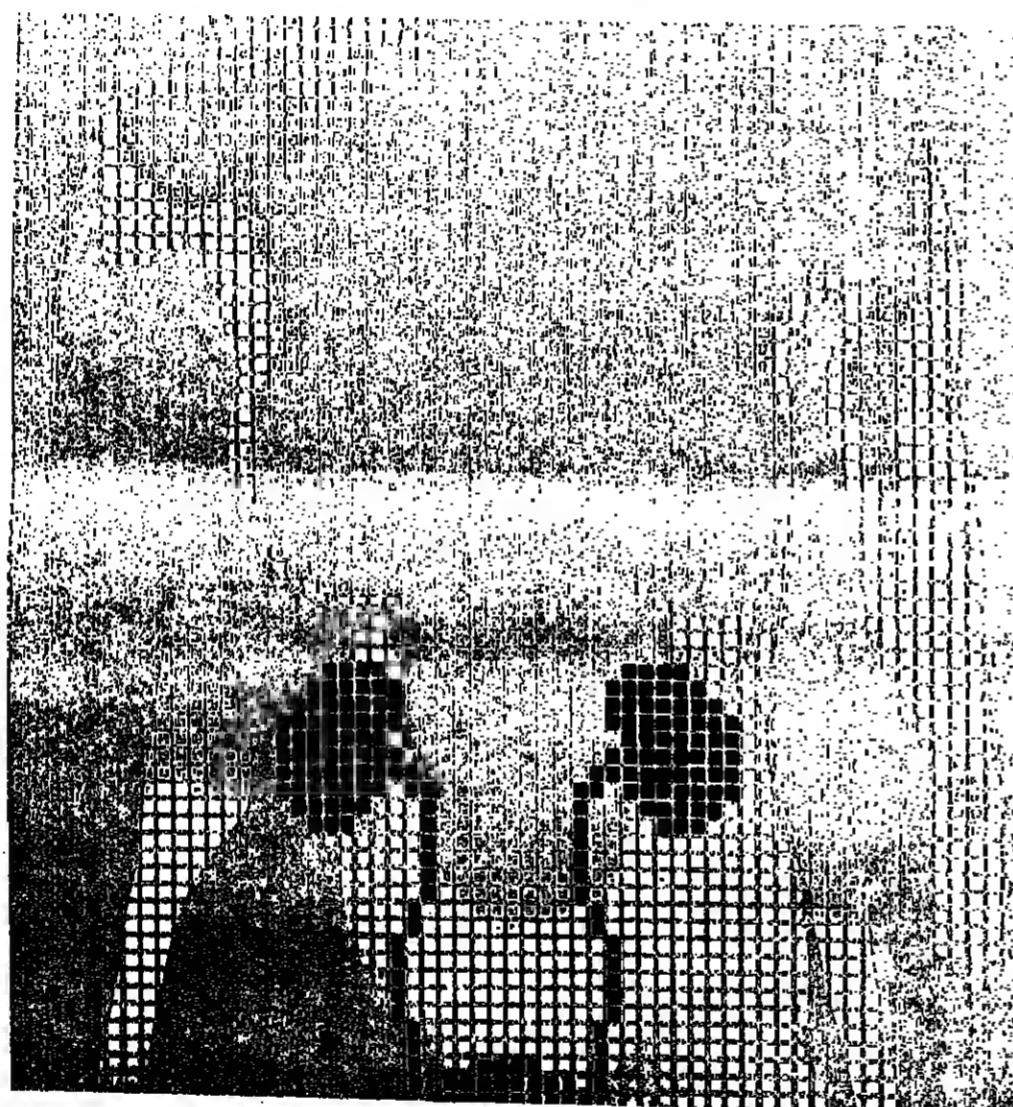


Azo

(Each tablet contains 100 mg phenazopyridine HCl

for what she's aware of

The symptoms that brought her to you Urgency, frequency, burning—these are the discomforting symptoms of cystitis that caused the patient to seek your help. Lasting relief depends on controlling the infection. But immediate relief may call for an analgesic. This is the patient who needs Azo Gantanol®. Azo to relieve symptoms; the action of Gantanol® (sulfamethoxazole) to control the bladder infection.



Before prescribing, please consult complete product information, a summary of which follows.

Indications: In adults, urinary tract infections complicated by pain (primarily pyelonephritis, pyelitis and cystitis) due to susceptible organisms (usually *E. coli*, *Klebsiella-Aerobacter*, *Staphylococcus aureus*, *Proteus mirabilis*, and, less frequently, *Proteus vulgaris*). In the absence of obstructive uropathy or foreign bodies. **Important Note:** Carefully coordinate *in vitro* sulfonamide sensitivity tests with bacteriologic and clinical response. Add amino-benzoic acid to culture media for patients already taking sulfonamides. Increasing frequency of resistant organisms currently limits the usefulness of antibacterial agents. Blood levels should be measured in patients receiving sulfonamides for serious infections, since there may be wide variations with identical doses; 12 to 15 mg/100 ml is considered optimal for serious infections; 20 mg/100 ml should be the maximum total sulfonamide level, as adverse reactions occur more frequently above this level.

Contraindications: Children below age 12; sulfonamides hypersensitivity; pregnancy at term and during nursing period. Contraindicated in glomerulonephritis, severe hepatitis, uremia, and pyelonephritis of pregnancy with gastrointestinal disturbances, because of phenazopyridine HCl component.

Warnings: Safe use in pregnancy has not been established, and teratogenic potential has not been thoroughly investigated. Deaths from hypersensitivity to HCl component.

Precautions: Children below age 12; sulfonamides hypersensitivity; pregnancy at term and during nursing period. Contraindicated in glomerulonephritis, severe hepatitis, uremia, and pyelonephritis of pregnancy with gastrointestinal disturbances, because of phenazopyridine HCl component.

Adverse Reactions: **Blood dyscrasias:** agranulocytosis, aplastic anemia and other blood dyscrasias have been reported; clinical signs such as sore throat, fever, pallor, purpura or jaundice may be early indications of serious blood disorders. Complete blood counts and urinalysis with careful microscopic examination should be performed frequently during sulfonamide therapy.

Precautions: Use with caution in patients with impaired renal or hepatic function; severe allergy, bronchial asthma and in glucose-6-phosphate dehydrogenase-deficient individuals. In the latter, hemolysis, a frequently dose-related reaction, may occur. Maintain adequate fluid intake to prevent crystalline end stone formation.

Adverse Reactions: **Blood dyscrasias:** agranulocytosis, aplastic anemia, thrombocytopenia, leukopenia, hemolytic anemia, purpura, bypass thrombocytopenia and methemoglobinemia; **allergic reactions:** erythema multiforme, necrotic, urticaria, serum sickness, pruritis, exfoliative dermatitis, anaphylactoid reactions, phototoxicity, edema, conjunctival and scleral injection, photokeratolysis, arthralgia and allergic myocarditis; **gastrointestinal reactions:** nausea, cramps, abdominal pain, hepatitis, diarrhea, anorexia, pancreatitis and stomatitis; **C.N.S. reactions:** headache, peripheral neuritis, motor depression, convulsions, drowsiness, hallucinations, tinnitus, vertigo and insomnia; **adverse reactions:** drug fever, chills, toxic nephritis with oliguria and anuria, polyarteritis nodosa and L.E. phenomenon. Due to certain chemical similarities with some sulfonamides, diuretics (acetazolamide and thiazides)

Gantanol®

and 0.5 Gm sulfamethoxazole.)

for what you're aware of

Bacterial infection

In just 2 to 3 hours after the initial adult dose, therapeutic blood and urine levels begin fighting *E. coli*, as well as susceptible strains of *Klebsiella-Aerobacter*, *Staph. aureus*, *Proteus mirabilis* and, less frequently, *P. vulgaris*.

She'll feel better while she gets better

As the Gantanol (sulfamethoxazole) component begins to fight the infection, analgesic Azo starts to relieve symptoms associated with bladder inflammation and irritation. For symptomatic cystitis, prescribe Azo Gantanol to help your patient feel better while she gets better.

New Electronic Device Permits
'Electromapping' of Heart: Page 2

International Headache Symposium—
A Critique: 'Current Opinion,' Page 5

Medical Tribune

and
Medical News

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world news of medicine and its practice—fast, accurate, complete

Wednesday, September 29, 1971
Vol. 12, No. 38

Liver Toxicity Of Methotrexate Held Doubtful

Medical Tribune World Service

OSLO—Evidence that the unusual liver findings seen in psoriasis patients treated with methotrexate are related in the disease rather than to the drug was presented here at the 19th meeting of the Nordic Dermatological Society.

Dr. Hugh Zacharine, of Marselisborg Hospital in Aarhus, Denmark, described 69 liver biopsies performed in 47 patients with psoriasis who were on treatment with methotrexate or considered for treatment.

When liver biopsies from 20 patients investigated before treatment were compared with those from 25 treated psoriasis, no significant differences were found. A high incidence of pathologic findings was observed in both groups.

Patients with psoriatic erythroderma showed the highest frequency of abnormal liver findings. Nine of 10 had increased fatty infiltration, and eight of 10 displayed signs of focal necrosis. The findings were not correlated with abuse of alcohol, obesity, or methotrexate therapy. No cases of cirrhosis were seen.

It is suggested that pathologic liver biopsies in psoriasis may be related to the disease," he declared. "Long-lasting topical treatment or use of other drugs, however, could also contribute."

He said that 10 patients underwent two biopsies, three patients three biopsies, and two patients four biopsies, with an average interval of 10 months. These serial biopsies showed a mild tendency toward increased fatty acids. Patients treated for more than two years with methotrexate.

Continued on page 16

Chinese Medicine Is Old, New, Eastern, Western



Medical practice today in China combines Western techniques with traditional, centuries-old methods. Above, group of health students test acupuncture on themselves.

Barefoot Doctors First Link In Massive Health Program

Medical Tribune World Service

CANTON, CHINA—"Barefoot doctors" are bringing medical aid to the 600,000,000 workers and peasants of China as the first step to what could become the world's most massive health care system. They provide the first link in a chain of medical aid that reaches from tiny rural villages to major hospitals having full general and specialist staffs in Peking and other main cities.

Barefoot doctors are, in fact, neither barefoot nor doctors. A cross between a first-aid worker and a district nurse, they run both rural and industrial medical services with what officials describe as diligence and frugality. Their work has gone far towards solving a problem that troubles many other countries to one degree or another—how to provide remote areas with adequate medical service.

The exact number of barefoot doctors is not made available by the authorities. Guesses range between 1,000,000 and 5,000,000 throughout the country, and a story in a Peking newspaper mentioning that 12,000 of them function in the area of the capital suggests that this estimate is probably correct.

The method was applied to camptothecin—a naturally occurring anticancer compound of current interest—and some of its inactive derivatives by Robert F. Flury, Ph.D., Professor of Chemistry at Louisiana State University.

The method permitted an explanation

Exercise Plan To Evade CHD May Be Danger

Medical Tribune Report

JACKSON, Wyo.—Exercise programs for the prevention of coronary heart disease should be prescribed only as part of a broad preventive program, and stress electrocardiographic studies should always be carried out before exercise is recommended, the 31st annual Congress on Occupational Health of the American Medical Association was told here.

"Moderately strenuous exercise, taken as an isolated thing unto itself, probably provides no benefit and certainly exposes the patient to great risk," according to Dr. Brendan Phibbs, of the University of Arizona College of Medicine.

"There is no real point in forcing a tense, cigarette-smoking, overweight executive to jog a mile a day while he worries about the accountants and sees with resentment his life situation," he declared.

"You are likely to do much more harm than good; the man's circulating catecholamines are likely to reach astronomical levels; the diseased coronary artery that had not previously been manifested may find itself inadequate for the demands of the myocardium, and another death may be added to the already impressive total.

"On the other hand, if this executive

Continued on page 16

Method Checks Cancer Drug Camptothecin and Derivatives

Medical Tribune Report

WASHINGTON—A computerized method of screening anticancer compounds is both quicker and cheaper than current laboratory screening and should aid in designing completely new classes of active compounds, the American Chemical Society was told at its 162nd national meeting here.

The method was applied to camptothecin—a naturally occurring anticancer compound of current interest—and some of its inactive derivatives by Robert F. Flury, Ph.D., Professor of Chemistry at Louisiana State University.

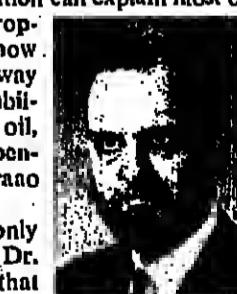
The method permitted an explanation

of the different activities of the compounds and a prediction of a new compound with activity similar to camptothecin, Dr. Flury said.

The method utilizes theoretical calculations that consider the distribution of electrons in molecules of the compound, he said. This distribution can explain most of the molecule's properties, including how reactive it is, the way it reacts, its solubility in water or oil, and its ability to penetrate the membrane of a cell.

There was only one property, Dr. Flury said, that could explain camptothecin's activity against cancer and its derivatives: inactivity—the value and direction of the dipole moment, a measure of unequal electron distribution in molecule. This property was significantly lower in camptothecin than in its derivatives, suggesting that camptothecin can penetrate more readily into the cell.

The recommendation was described as a "logical step, the next phase" in rubella control by Dr. H. Bruce Dull, executive secretary of the committee. While the vaccination of children will continue, more community resources may now be directed to the task of immunizing adolescent girls and adult women.



Dr. Flury

PHS Urges Rubella Vaccine Extension

Medical Tribune Report

ATLANTA, Ga.—The desirability of extending programs of rubella vaccination in adolescent girls and adult women was emphasized in a revised recommendation by the United States Public Health Service Advisory Committee on Immunization Practices.

"Because of the precautions which must apply, potential vaccinees in these groups should be considered individually," the committee pointed out. "They should receive vaccine only if they are shown to be susceptible by serologic testing, and if they agree to prevent preg-

nancy for two months after immunization."

The committee added: "To accomplish such extended use of rubella vaccine, the value and direction of the dipole moment, a measure of unequal electron distribution in molecule. This property was significantly lower in camptothecin than in its derivatives, suggesting that camptothecin can penetrate more readily into the cell.

A new compound, consisting of camptothecin molecules with one segment removed, was calculated to have close to the same properties as camptothecin itself and is predicted to have similar activity. It is currently being synthesized by a cooperator.

Continued on page 20

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(Each tablet contains 0.5 Gm sulfamethoxazole and 100 mg phenazopyridine HCl.)

**B.I.D. therapy
for the symptoms, for the infection**

Roche Laboratories
Division of Hoffmann-La Roche Inc.
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Continued on page 16

Electromap Gauges Thoracic Areas, Converts Data to Electric Impulses

Medical Tribune World Service

MILAN, ITALY—A new electronic device that its inventors claim is opening "new horizons" in heart disease research has been developed here by the Sime's Institute of Experimental Cardiology.

The device automatically measures 240 points of the thoracic surface and converts the information every two milliseconds into electric impulses. An on-line computer translates the information into figures or directly prints an "electromap" of the heart.

Prof. Bruno Taccardi, director of the electrophysiologic laboratory of the institute, said that experiments in both animals and human beings have proved the value of the device.

"Recent investigations carried out in this laboratory and in other research centers have shown that electromaps, which illustrate the instantaneous distribution of cardiac potentials on the body surface, yield more information on the electrical activity of the heart than can be obtained from the conventional electrocardiogram," he said.

Clinical Applications Limited

Although clinical applications of the device are still limited, Prof. Taccardi said, chest maps have provided information on the electric activity of the heart, he said, and is expected to solve certain problems related to final clinical applications of the new method.

The study, to be conducted in cooperation with various universities and hospitals, will produce much more precise information on the electric activity of the heart, he said, and is expected to solve certain problems related to final clinical applications of the new method.

As part of the method, electrocardiograms are recorded on the anterior and posterior surface of the trunk in groups of four tracings at a time, using a multibeam cathode-ray oscilloscope. A reference electrocardiogram, generally the VR lead, is recorded, together with each group of chest electrocardiograms.

The amplitude of each tracing is measured during ventricular activation and recovery. The amplitudes are converted into potential values and plotted on maps of the area explored. A map is thus built up drawn by equipotential lines.

Professor Taccardi noted that in heart disease a normal electrocardiogram sometimes leads to diagnostic problems and a

Medical Fees in New Guinea High at Two Pigs a Visit

Medical Tribune World Service

SYDNEY, AUSTRALIA—Churchmen and medical workers in New Guinea have called for a thorough investigation of the practice of witchcraft on the island, it was reported here. Missionaries say that more sorcery is practiced there than modern medicine.

So strong is the grip of the witch doctors that they often accompany patients into government hospitals to continue their treatment. Fees are high—as much as two pigs per consultation, according to the natives.

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Cancer Research in U.S.S.R.



WHO Photo

Investigators, above, at Cancer Research Institute, Moscow, study the immunology of spontaneous tumors occurring in dogs. At nearby Gamaleya Institute, a group of researchers have made significant contributions to knowledge of cancer of the liver.

Phlebography Found to Pinpoint Thrombi Post Mortem

Medical Tribune World Service

COPENHAGEN—Postmortem clarification of the frequency, localization, and extension of thrombi in a geriatric clientele has been obtained by intravenous phlebography with contrast medium injected into the calcaneus.

Dr. Lennart Diener, of the radiologic department of the Karolinska Hospital, Stockholm, told the 13th annual meeting of the International College of Angiology here that the method could possibly replace dissection to some degree and complement autopsy in more complete studies of the leg veins.

"The method can also be of interest in countries which are more restrictive concerning autopsy," he added.

The body is placed on a tilt table and fastened with a belt across the chest, he explained. The body is then brought to a semivertical foot-down position so that the heavy contrast medium can fill the veins from below without layering phenomena.

As contrast medium, Dr. Diener uses 0.1 Gm. barium sulfate per ml. of suspension. When injected into the calcaneus, it passes directly into the deep veins of the leg without filling the superficial venous system.

He reported having performed post-mortem phlebography as well as complete dissection of leg veins to the malleoli in 400 autopsies.

By increasing the amount of injected contrast medium, the phlebography could

be extended to include the caval system and the abdominal and possibly thoracic vessels, said Dr. Diener.

Japan Government Reimburses For Pollution-Linked Ailments

Medical Tribune World Service

TOKYO—The Japanese Government has appropriated 74,370,000 yen (\$206,580) as compensation for persons suffering from pollution-related ailments in 1971.

Buichi Oishi, director general of the Environmental Agency, said that 3,993 persons are being aided this year and that the agency is planning a stepped-up aid budget next year.

Currently, patients whose ailments have been traced to air pollution receive a 2,000-yen (\$5.55) allowance for commuting to hospital between three and 15 days a month and an allowance of 3,000 yen (\$8.33) if the hospital trips take up more than 15 days a month. Similar allowances are made to victims of water pollution.

Meanwhile, authorities in the industrial city of Tokai, south of Nagoya, have begun a program of free medical care for all patients suffering from pollution-inspired respiratory diseases. The measure is the first of its kind ever instituted in Japan and covers patients suffering from chronic bronchitis, bronchial asthma, asthmatic bronchitis, and pulmonary emphysema.

Commenting on these findings, Dr. Guthe said it is likely that mass screening, however cumbersome, will probably have to be employed more widely in the future. He noted that automation of the Treponema pallidum hemagglutination test appears to show promise.

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Year-Old Kansas Abortion Law Only 2% Loss Seen With New Technique Of Red Cell Thawing

Medical Tribune Report

Washington Bureau

pregnancy, and the majority of the women undergoing the operations were 18 to 22 years old, unmarried (73.6 per cent), and white (89.4 per cent), and spent one day in the hospital.

Statistics Not Alarming

Dr. Edwin Lyman, executive director of the Board of Health, said that for the most part the statistics were not alarming nor did they seem out of line.

"I have nothing to compare with," Dr. Lyman commented. "This is our first year of experience. I can state that the law has not altered significantly the state's birth rate—there was no abrupt drop in births. This indicates to me that the number of abortions that used to be carried out illegally in the state just about equaled the number carried out legally last year."

With regard to the fact that more out-of-state residents (5,315) obtained abortions than Kansans (3,244), Dr. Lyman observed that the majority of the operations were performed in Kansas City, Kansas, which is just across the street from Kansas City, Missouri, where abortions are still illegal under most circumstances.

The Board of Health turned over the matter to the State Attorney General's office for investigation. Under the abortion statute, violators could be charged with a class D felony, which carries a sentence of one to 10 years in the state penitentiary and a fine up to \$5,000.

Meanwhile, some Kansas physicians and legislators expressed concern over the first year's abortion tally, which they said was twice as high as was predicted when the law was passed. Dr. Francis Bice of Wakeeney, health board member, commented that the "sickening" total should show legislators "how much a bunch of damn fools they were to pass such a law."

10-Day Ban Declared

At the University of Kansas Medical Center, Dr. William O. Rieke, who became vice-chancellor of medical affairs in July, declared a 10-day ban on abortions August 20. He explained that it was in accordance with a general hospital directive limiting all elective surgery procedures through August 30 because of a nursing shortage.

Dr. Rieke admitted in an interview, however, that the ban was also instituted so that he could "verify that all our procedures are done under the best conditions."

He noted that therapeutic abortion is an emotional political issue in Kansas at this time and added: "I don't think people appreciate the hot spot I'm sitting in." The medical center staff performed about 2,200 of the state's therapeutic abortions last year.

The medical center's administration already had decided to omit any reference to abortion in the Hippocratic Oath administered to medical students graduating this year.

After a special meeting with Dr. Kermit Krantz, chief of the obstetrics and gynecology department, and other department personnel, Dr. Rieke issued a statement saying, "I have every assurance that the letter and spirit of Kansas law have been met in the past and will be in the future," and removed the general ban on abortions after August 30.

According to the first year's statistics on Kansas therapeutic abortions, 88.8 per cent were performed to preserve the mother's mental health; 85.8 per cent were performed during the first trimester of

pregnancy, and the majority of the women

undergoing the operations were 18 to 22

years old, unmarried (73.6 per cent), and

white (89.4 per cent), and spent one day

in the hospital.

Contingentous Hypersensitivity to sulfonamides, infants less than 2 months of age, pregnancy at term, and during the nursing period.

Indications: Acute, recurrent or chronic urinary tract infections (primarily cystitis, pyelitis, pyelonephritis) due to susceptible organisms (usually *E. coli*, *Klebsiella*, *Aerobacter*, *Staphylococcus aureus*, *Pseudomonas aeruginosa*) and test tube sensitivity to sulfonamides.

Contraindications: Patients with a history of hypersensitivity reactions, anuria, oliguria, anuria, anuria, and other blood dyscrasias associated with sulfonamides, adrenalin, chloramphenicol, and other drugs.

Precautions: Patients with certain chronic diseases, particularly those with renal, hepatic, and bone marrow dysfunction, should be monitored closely.

Adverse Reactions: Blood dyscrasias, agranulocytosis, aplastic anemia, and other blood dyscrasias, particularly those associated with sulfonamides.

Supplied: Tablets containing 0.5 Gm. sulfisoxazole.

by passing a glucose-saline solution through the red cells while they are being centrifuged. The process takes about 30 minutes.

Dr. Meryman said that a freezing method developed five years ago by Dr. Charles Higgins, of Massachusetts General Hospital, causes a loss of about 25 per cent of the red blood cells.

Red Cross officials predicted that by the end of this year 50,000 units of frozen washed blood will be available and that 18 regional centers will be in a position to process 100,000 units of frozen blood.

The cost of processing the frozen cells, \$37 a pint, compared with \$12 for regular donated blood, will be a limiting factor in their availability, they noted.

In addition to the Boston and Washington, D.C., centers, the A.R.C. is sending freezing machinery to blood banks in Atlanta, Ga.; Birmingham, Ala.; Nashville, Tenn.; Charlotte, N.C.; Cleveland; Columbus, Ohio; Detroit; St. Louis; Albany and Rochester, N.Y.; Los Angeles; Portland, Ore.; Tucson, Ariz.; St. Paul; Lansing, Mich.; and Hartford, Conn.

When the problem

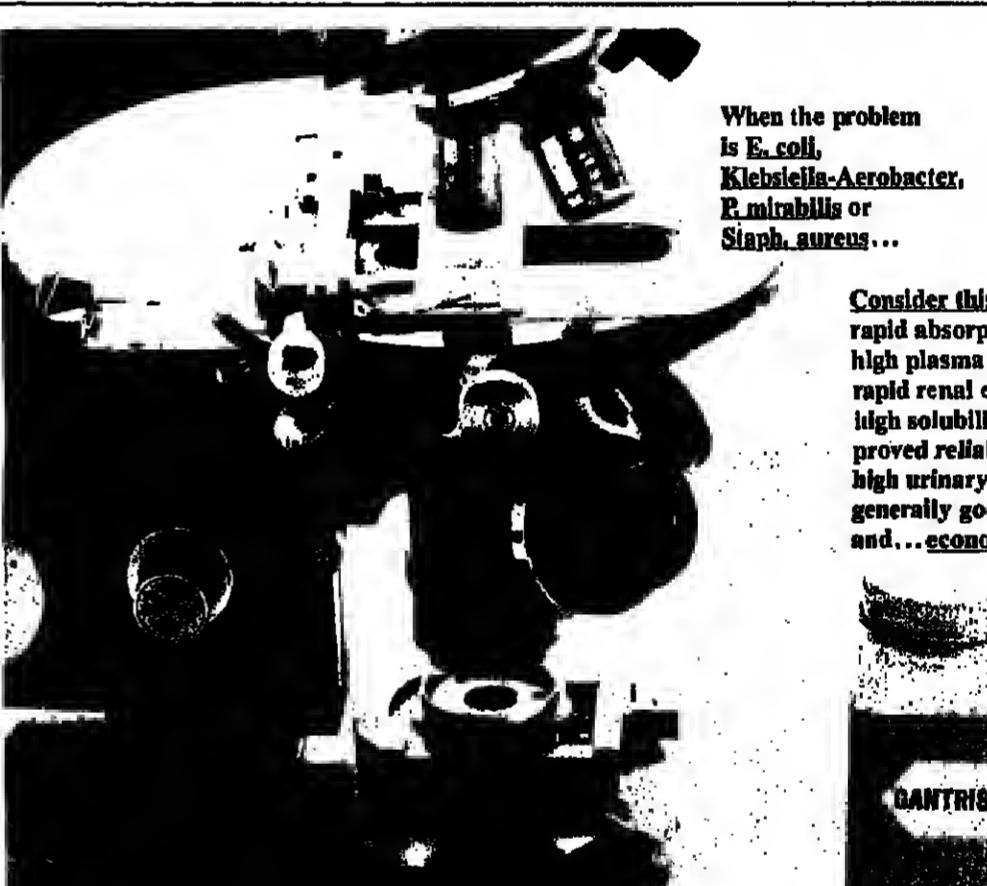
is *E. coli*,

Klebsiella-Aerobacter,

P. mirabilis or

Staph. aureus...

Consider this:
rapid absorption,
high plasma concentrations,
rapid renal clearance,
high solubility at urinary pH,
proved reliability,
high urinary drug levels,
generally good tolerance,
and...economy.



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sulfisoxazole/Roche®

classic for nonobstructed cystitis,

pyelitis and pyelonephritis

4 to 8 tablets stat, 2 to 4 tablets q.d.

Before prescribing, please consult complete product information, a summary of which follows.

Indications: Acute, recurrent or chronic urinary tract infections (primarily cystitis, pyelitis, pyelonephritis) due to susceptible organisms (usually *E. coli*, *Klebsiella*, *Aerobacter*, *Staphylococcus aureus*, *Pseudomonas aeruginosa*) and test tube sensitivity to sulfonamides.

Contraindications: Patients with a history of hypersensitivity reactions, anuria, oliguria, anuria, and other blood dyscrasias associated with sulfonamides.

Precautions: Patients with certain chronic diseases, particularly those with renal, hepatic, and bone marrow dysfunction, should be monitored closely.

Adverse Reactions: Blood dyscrasias, agranulocytosis, aplastic anemia, thrombocytopenia, leukopenia, hemolytic

anemia, purpura, hypoprothrombinemia, methemoglobinemia. **Allergic reactions:** Erythema multiforme (Stevens-Johnson syndrome), generalized skin eruptions, epidermal necrolysis, urticaria, serum sickness, pruritis, scabies, scrotal dermatitis, impetigo, cellulitis, cellulitis, and other forms of cutaneous and scleral infections, phlebitis, vasculitis, the presence of autoimmune or foreign bodies. Important: *In vitro* sulfonamide sensitivity tests are not always reliable; tests must be coordinated with bacteriologic and clinical response. Ambien-like side effects may be early indicators of serious blood disorders. **Diagnosis:** Clinical blood counts and urinalysis with careful microscopic examination should be performed frequently during sulfonamide therapy.

Precautions: Patients with certain chronic diseases, particularly those with renal, hepatic, and bone marrow dysfunction, should be monitored closely.

Adverse Reactions: Blood dyscrasias, agranulocytosis, aplastic anemia, thrombocytopenia, leukopenia, hemolytic

Outpatient Care on Rise

Clip and for every patient admitted to a hospital in 1970, there were 5.7 outpatient visits, reports the American Hospital Association. Ambulatory care continues to be the fastest-growing service in the nation's hospitals.

In 1970 the 7,123 hospitals registered by the American Hospital Association reported a total of 181,400,000 outpatient visits, for an increase of 11.1 per cent over 1969.



all in the same boat many hypertensives do better on **Esimil**

guanethidine monosulfate 10 mg
hydrochlorothiazide 25 mg

Esimil may get things moving for all sorts of patients with moderate to severe hypertension. Often controls blood pressure when other antihypertensives fail. Usually keeps it controlled, too.

The key is guanethidine—perhaps the most effective antihypertensive available.

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Esimil®
guanethidine monosulfate 10 mg
hydrochlorothiazide 25 mg

Indications:

Hypertension (other than labile forms) which cannot be adequately controlled with simpler agents; hypertension almost all forms of fixed and progressive hypertension disease, when side effects of other antihypertensives prevent effective treatment.

Contraindications:

Hyperthyroidism or suspected pheochromocytoma; hypersensitivity to guanethidine. Do not use with MAO inhibitors.

Hydrochlorothiazide Anuria, discontinuous drug if renal shutdown occurs for any reason. Progressive development of hepatic coma. Do not give to patients known allergic to thiazides or other sulfonamide-derived drugs.

Warnings:

Guanethidine and hydrochlorothiazide are potential clinical problems. Physicians should be familiar with both drugs and their combination before prescribing, and patients should be warned not to discontinue therapy.

Guanethidine: Warnings about the potential hazards of orthostatic hypotension, which can occur frequently. To prevent fainting, patients should sit or lie down when onset of dizziness or orthostatic hypotension occurs. If orthostatic hypotension during initial dosage adjustment and with postural changes. Postural hypotension is most marked in the morning and is accounted for by orthostatic hypotension. Patients should be warned to avoid sudden or prolonged standing or sitting while taking guanethidine.

Concurrent use with tricyclic derivatives may cause orthostatic hypotension, and may lead to tubocurarine. If possible, withdraw therapy 2 weeks prior to surgery. Hypotensive episodes may have been observed. If emergency surgery is indicated, antihypnotic and anesthetic agents should be administered in reduced dosage. Give oxygen, atropine, and vasodilators ready for immediate use. Give vasodilators with caution to patients with a history of bronchial asthma.

Guanethidine: The safety of guanethidine for use in pregnancy has not been established. Therefore, it should not be used in pregnant patients or women of childbearing age. If the physician, it is deemed essential to the welfare of the patient.

Hydrochlorothiazide: Thiazides should be used with caution in pregnant or lactating patients. It is required when treating patients with hypertension of the pregnant or lactating woman.

bronchial asthma, since the condition may be aggravated.

Hydrochlorothiazide: Small bowel stenosis, with or without ulceration, has been reported with use of enteric-coated thiazides with potassium, and with enteric-coated potassium alone. These bowel lesions have caused obstruction, hemorrhage, and death. These have been reported and death has occurred. Although the incidence of these lesions is low, and a causal relationship in man has not been definitely established, enteric-coated potassium should not be implicated. Therefore, enteric-coated potassium containing formulations should be used only when dietary supplementation is not practical and discontinued when possible.

Lowering of blood pressure in hypertensive pa-

tients may sometimes result in nitrogen retention, to renal blood flow reduced, particularly in those with preexisting renal disease. Prolonged renal insufficiency is often a disabling side effect of thiazides. Hypotension may develop in those with impaired renal function. Dose reduction always be carefully titrated.

Thiazides may decrease arterial responsiveness to norepinephrine and increase responsiveness to tubocurarine. If possible, withdraw therapy 2 weeks prior to surgery. Hypotensive episodes may have been observed. If emergency surgery is indicated, antihypnotic and anesthetic agents should be administered in reduced dosage. Give oxygen, atropine, and vasodilators ready for immediate use. Give vasodilators with caution to patients with a history of bronchial asthma.

Guanethidine: The safety of guanethidine for use in pregnancy has not been established. Therefore, it should not be used in pregnant patients or women of childbearing age. If the physician, it is deemed essential to the welfare of the patient.

Hydrochlorothiazide: Thiazides should be used with caution in pregnant or lactating patients. It is required when treating patients with hypertension of the pregnant or lactating woman.

diabetic patients may be increased, decreased, or unchanged. Lactic dehydrogenase may become elevated during thiazide therapy.

If nitrogen retention indicates onset of renal impairment, discontinue drug.

Adverse Reactions:

Guanethidine: Give cautiously to patients with severe coronary insufficiency, recent myocardial infarction, or cerebrovascular insufficiency. Give slowly. Use extreme caution in those with severe cardiac failure.

Appetite suppressants (eg, amphetamine), mild stimulants (eg, ephedrine, mephedrine), and tricyclic antidepressants (eg, imipramine, propantheline, doxepin, mianserin) may have an additive effect of guanethidine. Waller was after discontinuing MAO inhibitors before starting guanethidine.

Guanethidine: Severe reactions due to sympathetic blockade—blushing, sweating, fainting, syncope. Frequent reactions caused by unopposed parasympathetic activity—bradycardia, increase in bowel movements, diarrhea (which may be profuse and rectal bleeding may occur with the drug). Other common reactions—Inhibition of ejaculation, fluid retention, edema, congestive heart failure. Less frequently—dyspnea, fatigue, nausea, vomiting, headache, dizziness, tachycardia, dizziness, scalding heat, dry mouth, rise in BUN, rise of the lido, blurring of vision, periorbital edema, or other chronic disorders may be aggravated by a relative increase in parasympathetic tone. Periodic blood counts and liver function tests are advised during prolonged therapy.

Hydrochlorothiazide: Perform serum potassium, BUN, uric acid, blood urea nitrogen, and creatinine at appropriate intervals during therapy. Watch patients for clinical signs of fluid or electrolyte imbalance (hypotension, hypochlorhydria, hypokalemia, headache, xanthopsia, decreased urine output, etc). Watch for signs of dryness of mouth, thirst, weakness, tachycardia, hypotension, muscle cramps or cramps, muscular fatigue, hypotension, oliguria, tachycardia, GI disturbances, hypotension, and/or electrolyte abnormalities. Use cautiously in diabetics. Hypoglycemia may occur but is generally reversed by a glucose agent.

Thiazides may decrease arterial responsiveness to norepinephrine and increase responsiveness to tubocurarine. If possible, withdraw therapy 2 weeks prior to surgery. Hypotensive episodes may have been observed. If emergency surgery is indicated, antihypnotic and anesthetic agents should be administered in reduced dosage.

The possibility of sensitivity reaction should be determined. If sensitivity reaction occurs, discontinue thiazide with a history of allergy or bronchial asthma.

Guanethidine: The safety of guanethidine for use in pregnancy has not been established. Therefore, it should not be used in pregnant patients or women of childbearing age. If the physician, it is deemed essential to the welfare of the patient.

Hydrochlorothiazide: Thiazides should be used with caution in pregnant or lactating patients. It is required when treating patients with hypertension of the pregnant or lactating woman.

International Headache Symposium

By DONALD J. DALESSIO, M.D.

Head, Division of Neurology, Scripps Clinic and Research Foundation, La Jolla, Calif.

THE AUTHOR recently returned from the International Headache Symposium held in Elsinore, Denmark, should confess that he was one of the organizers of the conference—lucky what follows cannot be said to be an entirely objective critique.

Several entirely new observations were reported at the meeting. G. W. Bruyn of

the Netherlands found that patients with migraine have elevated blood ammonia levels of significance. In 12 normal controls he found that the plasma ammonia ranged from 21 to 63 micrograms/100 ml, with a mean of 40 micrograms/100 ml. In 22 patients with migraine, the levels ranged from 39 to 390 micrograms/100 ml, with a mean of 103 micrograms/100 ml. These patients did not have liver disease. This important observation requires confirmation and further biochemical investigation, but it may provide another clue to a dietary abnormality producing headache, related to a resting hyperammonemia. Should this prove to be the case, some form of protein restriction in the diet might be advisable for susceptible patients with migraine.

Treatment of rats with methysergide produced a protective influence against the serotonin-depleting activity of reserpine. Similar protection could not be produced with ergotamine. Lance of Australia has demonstrated reduced platelet serotonin levels prior to a migraine episode. Thus the protective effects of methysergide mentioned above may relate to its use in the prophylaxis of migraine.

Several physiologic studies were pre-

pared which demonstrated again that in the prodromal phase of migraine, a reduction in cerebral blood flow occurs, at times to levels critical for adequate oxygenation of the brain. Skinhøj of Copenhagen, using intracarotid radioactive xenon injection for determination of regional cere-

bral blood flow, has found that there is diminished local cerebral blood flow of great significance in the prodromal phase of migraine. Thereafter, during the headache phase, a luxury perfusion state occurs, associated with intracerebral edema, as would be expected if the initial cerebral ischemia were severe. O'Brien of Newcastle upon Tyne, England, finds that the changes measured by radioactive techniques in the prodromal phase of migraine are widespread and bilateral, and not necessarily related to localized aura symptoms.

G. S. Barolin of Austria, presenting the Harold G. Wolff Lecture, reviewed his experience with electroencephalographic findings in migraine. F. G. Silman of Jerusalem offered the first paper in this author's recent memory relating disease state to climate. The hot, dry winds of Israel may produce headache in sensitive subjects. In 70 of 80 patients so affected, there was an increased urinary serotonin excretion associated with the heat stress.

What was new in therapeutics? The English reported on their further trials with clonidine, an antihypertensive drug which also reduces vasoconstrictor activity and which shows promise of being a useful agent in migraine prophylaxis. Dr. Marshall Wilkin of London presented data to show that persons subject to headache after ingestion of tyramine are most likely

to benefit from chronic therapy with clonidine. Also, several papers from different areas of Europe and Israel reported on the prophylactic effects of BC-1015, which has proved effective in reducing the frequency and intensity of vascular headache and has some antidepressant properties.

Papers on clinical pharmacology were lacking from the large U.S. group present. It seems evident that clinical research on new drugs in this country is fast becoming an obsolete and occult pursuit. One might observe that we have come full cycle and that pressures from the Food and Drug Administration, combined with the ever-present threat of malpractice suits, have served to make clinical pharmacology an exercise for those in other fields, who practice medicine in less antagonistic circumstances.

1962 was a bad year for Staph infection

In a hospital study from 1960 to 1967, only 2 positive staphylococcal lesions were noted among 34,262 infants washed with pHisoHex. Both occurred in 1962.

In 6 additional studies, * of 1974 infants washed with pHisoHex, Staph colonization was only 2.4%; while in 1960 "unwashed" control infants, colonization amounted to 45%.

Anti-Staph colonization for the infant usually begins with a pHisoHex bath before he is born. This can be continued and strengthened throughout the infant's stay in the hospital nursery by bathing him daily with pHisoHex and having everyone who handles the infant wash his hands with pHisoHex before and after handling the baby. This routine builds a cumulative, rinse-resistant film of anti-bacterial hexachlorophene on the skin to form a powerful barrier against Staph and many other bacteria.

Mother can maintain this anti-bacterial colonization at home by bathing baby exclusively with pHisoHex. And nonalkaline, hypoallergenic pHisoHex is kind to skin.

References: 1. Gluck, Louis: *Home Practice* 9:33, Jan. 1968 (author's correction). 2. Payne, Margaret C.; Wood, H. F.; Karkhoff, Walter, and Gluck, Louis: *Am. J. Epidemiol.* 82:305, Nov. 1965. 3. Gluck, Louis, and Wood, H. F.: *Arch. Epidemiol. J. Med.* 265:1177, Dec. 14, 1961. 4. Simon, H. J.: *Arch. Pediatr. and Adolesc. Med.* 108:254, Feb. 1965. 5. Gluck, Louis, and Wood, H. F.: *New England J. Med.* 265:1171, Dec. 14, 1961.

Winthrop Laboratories, New York, N.Y. 10018 *Winthrop*



CIBA

COMING NEXT ISSUE

• Marijuana

Survey finds 15 per cent of teen-agers have used "pot."

• Breast cancer

Careful screening may spare women from radical mastectomy.

• Vitamin deficiency

Debate on requirements may cloud extent of problem.

CIBA Pharmaceutical Company

Summit, New Jersey

Anxiety—frequent cause of cardiac complaints or concomitant of cardiac symptoms. Cardiac complaints may be signals of underlying anxiety or of organic cardiac disease. In either case, anxiety of varying severity often plays a prominent role in provoking functional complaints or complicating organic disease. Thus, whenever excessive anxiety is a significant component of the clinical profile, adjunctive use of Librium (chlordiazepoxide HCl) may be of value.

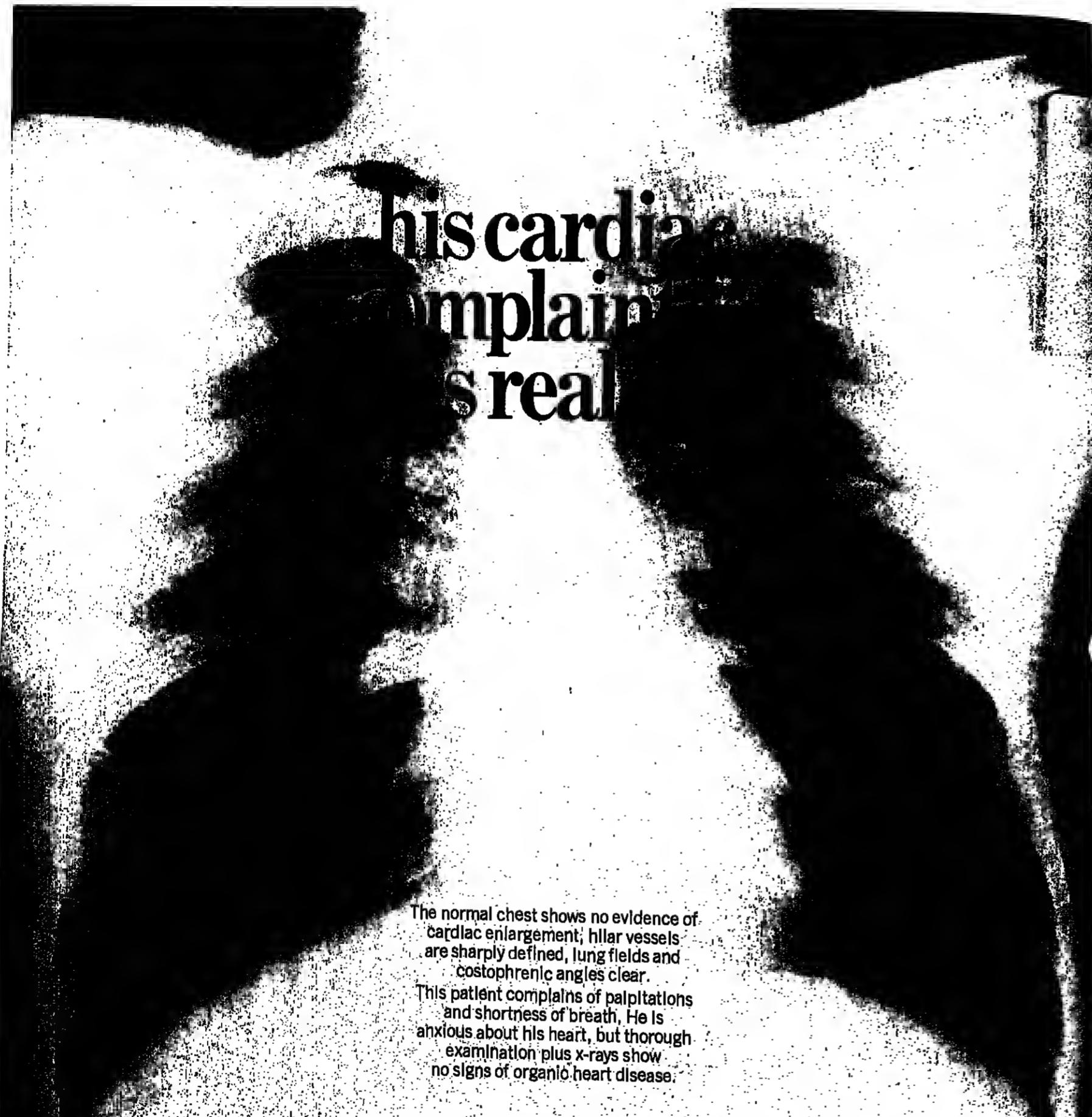
Librium (chlordiazepoxide HCl) is used concomitantly with certain specific medications of other classes of drugs, such as cardiac glycosides, diuretics and antihypertensive agents.

Librium (chlordiazepoxide HCl) is especially well suited for extended use because of its wide margin of safety. In general use, the most common side effects reported have been drowsiness, ataxia and confusion, particularly in the elderly and debilitated. (See summary of prescribing information.) Moreover, the anti-anxiety benefits of Librium are

generally maintained without diminution of effect or need for increase in dosage. When treatment is prolonged, periodic blood counts and liver function tests are advisable. **Three dosage strengths are available** for flexible, individualized therapy of various degrees of excessive anxiety in patients with functional or organic heart disorders.

ROCHE

Roche Laboratories
Division of Hoffmann-La Roche Inc.
Nutley, N.J. 07110



The normal chest shows no evidence of cardiac enlargement; hilar vessels are sharply defined, lung fields and costophrenic angles clear.

This patient complains of palpitations and shortness of breath. He is anxious about his heart, but thorough examination plus x-rays show no signs of organic heart disease.

Before prescribing, please consult complete product information, a summary of which follows:

Indications: Indicated when anxiety, tension and apprehension are significant components of the clinical profile.

Contraindications: Patients with known hypersensitivity to the drug.

Warnings: Caution patients about possible combined effects with alcohol and other CNS depressants. As with all CNS-acting drugs, caution patients against hazardous occupations requiring

complete mental alertness (e.g., operating machinery, driving). Though physical and psychological dependence have rarely been reported on recommended doses, use caution in administering to addiction-prone individuals or those who might increase dosage; withdrawal symptoms (including convulsions), following discontinuation of the drug and similar to those seen with barbiturates, have been reported. Use of any drug in pregnancy, lactation, or in women of childbearing age requires that its potential benefits be weighed

against its possible hazards. **Precautions:** In the elderly and debilitated, and in children over six, limit to smallest effective dosage (initially 10 mg or less per day) to preclude ataxia or oversedation. Increasing gradually as needed and tolerated. Not recommended in children under six. Though generally not recommended, if combination therapy with other psychotropics seems indicated, carefully consider individual pharmacologic effects, particularly in use of potentially drugs such as MAO inhibitors and

phenothiazines. Observe usual precautions in presence of impaired renal or hepatic function. Paradoxical reactions (e.g., excitement, stimulation and delirium) have been reported in psychiatric patients and hyperactive aggressive children. Employ usual precautions in treatment of anxiety states with evidence of impending depression; suicidal tendencies may be present and protective measures necessary. Variable causal relationship has not been established clinically.

Adverse Reactions: Drowsiness, ataxia and confusion may occur, especially in the elderly and debilitated. These are reversible in most instances by proper dosage adjustment, but are also occasionally observed at the lower dosage ranges. In a few instances syncope has been reported. Also encountered are isolated instances of skin eruptions, edema, minor menstrual irregularities, nausea and constipation, extrapyramidal symptoms, increased and decreased

libido—all infrequent and generally controlled with dosage reduction; changes in EEG patterns (low-voltage fast activity) may appear during and after treatment; blood dyscrasias (including agranulocytosis), jaundice and hepatic dysfunction have been reported occasionally, making periodic blood counts and liver function tests advisable during protracted therapy. **Supplied:** Capsules containing 5 mg, 10 mg or 25 mg chlordiazepoxide HCl.

For relief of moderate to severe anxiety associated with cardiovascular symptoms adjunctive

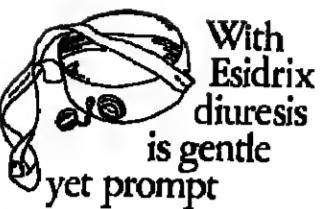
Librium® 10 mg
(chlordiazepoxide HCl)
1 or 2 capsules
t.i.d./q.i.d.



The heart is enlarged, the hilar vessels are engorged, and the bronchovascular markings accentuated. There are bilateral pleural effusions.

This patient complains of the same symptoms—palpitations and shortness of breath. He is concerned that he may have heart disease, and thorough examination plus x-rays justify his high level of anxiety.

Avoid the complications of "diuretic overdry"



No one denies there's a time and place for a highly potent natriuretic.

But most patients rarely need it. Which is why hydrochlorothiazide—originated by

CIBA as Esidrix—remains the most widely used oral diuretic.

With Esidrix you usually avoid the abrupt flushing out common with fast-acting natriuretics.

Diuresis is prompt; edema is relieved gradually over a 12-hour period. Which is usually fast enough. Just as important, it's smooth and gentle.

Things are complicated enough for the edema patient. Rely on Esidrix, the smooth, gentle diuretic. Particularly in maintenance therapy.

Moreover, the risk of serious salt and water loss is reduced.

However, since

fluid and electrolyte imbalance may occur, patients should be watched closely for clinical signs (please see brief prescribing information).

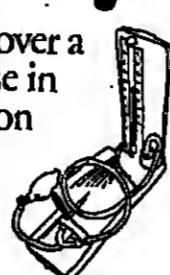
Esidrix is still unsurpassed as a diuretic-antihypertensive. Labeling for one newer natriuretic states:

"Hypertensive patients who cannot be adequately controlled with

Proven by over a decade's use in hypertension

thiazides will probably also not be adequately controllable with [furosemide] alone."

And Esidrix is amply proven alone in mild hypertension. As an adjunct in mild to severe cases.



Esidrix® (hydrochlorothiazide) is often just enough.

Dorothy Lamour—
star of stage, screen and television.

Esidrix® (hydrochlorothiazide)

Indications: Edema and hypertension. Contraindications: Anuria; discontinuous drug if renal shutdown occurs for any reason. Prostaglandin-induced edema may accelerate development of hepatic coma. Do not give to patients with known allergy to thiazides or to ceftriaxone, derived drugs.

Warnings: Small bowel stricture, with or without ulceration, has been associated with use of enantiomeric thiazides with potassium, and with enantiomeric potassium and potassium excretion. Thiazides may decrease glucose tolerance; use cautiously in diabetes. Hyperglycemia may occur, but is generally reversed by discontinuation.

Thiazides may decrease arterial responsiveness to noradrenergic and increased responsiveness to vasoconstrictors. If possible, withdraw therapy 2 weeks prior to surgery. Hypotensive episodes under anesthesia have been observed. If emergency surgery is indicated, preanesthetic and anesthetic agents should be administered in reduced dosage.

The possibility of sensitivity reactions should be considered in patients with a history of allergic or bronchial asthma. Use in Pregnancy: Thiazides should be used with caution in pregnant or lactating patients since this drug crosses the placental barrier and should be used only if the milk and may result in fetal hypochlorhydria, thrombocytopenia, or edema may be present.

Thiazides may precipitate edema, cumulative effects may develop in those with impaired renal function. Dosage should always be carefully titrated.

For special attention to electrolyte balance of patients with severe hepatic insufficiency, in patients with cirrhosis and ascites, and for symptoms of impending hepatic coma, confusion, drowsiness, stupor, or for increased arterial ammonia concentration, serum and potassium excretion. Thiazides may decrease glucose tolerance; use cautiously in diabetes. Hyperglycemia may occur, but is generally reversed by discontinuation.

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De Motu Cordis Updated

HARVEY'S ORPHAN DISCOVERY was the circulation of the blood, but his classic book is titled *De Motu Cordis*—i.e., "motion of the heart." Necessarily relying on direct observation, Harvey studied the heart beating in "very little Fish...all the body of which is transparent," in chick embryos placed in warm water, and in mammalian dissections. That is at least one order of magnitude removed from the array of modern techniques applied to investigate the movements of the heart and blood.

The very names of current investigative methods are formidable—e.g., videoscintigraphy, radionuclide angiography, radiokymography, biplane cineradiography, echocardiography, kinectocardiography, apex cardiography, electrokymography, roentgenkymography. Some of the newer procedures appear most promising.

Radionuclide angiography, developed by Dr. Joseph P. Kriss and colleagues at the Division of Nuclear Medicine of Stanford University, is particularly intriguing. As far as the patient is concerned, all that is required is the intravenous injection of a few cubic centimeters of technetium-99m-pertechnetate. The course of the injected radionuclide through the great vessels and chambers of the heart is followed in precise detail by a scintillation camera and a variable time-lapse video-scanlscope, permitting display, replay, and photography of cardiovascular morphology.

As described by Dr. Kriss and his colleagues in a detailed article in the June issue of *Circulation*, the procedure has a very broad scope of diagnostic applicability in heart disease, including congenital heart and valvular lesions, ventricular and atrial aneurysms, ventricular hypertrophy and cardiomyopathies, and pericardial effusion. The method has permitted the quantitative assessment of such hemodynamic functions as cardiac output, chamber size, and blood flow as reflected by transit times; and quantitation appears feasible also of intracardiac shunts, myocardial contractility, and coronary blood flow. Contrast angiography and cardiac catheterization, which are widely employed in diagnostic study of the heart, are both "invasive" techniques with inconvenience and some hazard to the patient. The particular virtue of radionuclotide angiography is its simplicity in addition to its versatility.

Cardiac diagnosis has come a long way

from palpation, auscultation, and the long-familiar techniques of fluoroscopy and electrocardiography, although, combined with good history, these still suffice in most patients. Harvey would doubtless have been much intrigued to see how far *De Motu Cordis* has come since his observations on transparent "little Fish."

Congratulations to the FDA

THE FOOD AND DRUG ADMINISTRATION should be commended for its choice of Peter Barton Hutt as chief counsel. Mr. Hutt comes to the agency with a decade of extensive experience in Food and Drug law, specializing in this area during his years with a leading Washington law firm as chairman of the American Bar Association's Food and Drug Law Committee, and as a member of the Advisory Lawyers Committee to the Food and Drug Law Institute. It is, however, not only technical excellence that Mr. Hutt brings to his new tasks but practicality, liberalism, and a concern for the public welfare. His years of working with Seniors and Representatives in the drafting of health-related legislation, mostly behind the scenes, have given him practical knowledge of the vagaries of politics. His concern with substance abuse has been long and lasting, ha

ving the first lawyer in the country to establish in the courts the principle that the alcoholic is sick and should not be the subject of criminal law. Lastly, and perhaps most important as he takes office, Mr. Hutt has always been a believer in the Constitutional guarantee of basic rights for the individual, the right of the doctor to practice medicine with pride and freedom, and the right of both to live and work without mortal fear of government interference and reprisal. Dr. Walter Modell once wrote, "There is also a strong possibility that in the future physicians will be frightened into using token dosage and treat patients in ways designed to assuage the FDA, rather than the disease." In commenting on this statement, Peter Barton Hutt wrote simply, "One would hope that this could be avoided."

We wish him well.

"Induced Traffic"

CHLINICAL QUOTE: "Another facet of autonomic sampling and analyzing of blood gases and pH can well be what traffic engineers refer to as 'induced traffic.' And if the hospital billing system were hooked into the same computer, the 349 tests could add approximately \$5,800 to the patient's bill." (Charles D. Cook, M.D., Automated Blood Analysis: Success or Excess? *Pediatrics* 48:1, July, 1971.)



"One leaspoonfull to be taken every full moon
—and kaop il out of reach of the children."

Diabetes Controversy—Continued

Editor, MEDICAL TRIBUNE:

It is my belief that the controversial report of the University Group Diabetes Program relative to the alleged acceleration of cardiovascular complications in adult-onset diabetes receiving tolbutamide—and presumably all related sulfonylurea compounds—can be better understood in the light of a basic issue. This concerns the role of the hyperinsulinized-diabetic state in the pathogenesis of ischemic (coronary) heart disease...

I have repeatedly stressed that the onset of angina pectoris or cardiac arrhythmias three or more hours after eating, especially during the night, is a hallmark of myocardial glycopenia, whether coronary occlusion is or is not demonstrable...

My review of the U.G.D.P. report warrants pointing out the following observations:

1. The greatest patient cooperativeness relative to taking 1.5 Gm. tolbutamide daily ("highest level of adherence to study medication") was in the tolbutamide-treated group. Specifically, 74 per cent in this group took all of their study medication for at least 75 per cent of the follow-up periods during which they were under study. The corresponding percentages for the placebo and insulin groups only ranged from 45.3 per cent to 69.8 per cent.

2. There was less hypoglycemia in the insulin-treated groups due to the unequivocal modification of the assigned dosage for those patients receiving tolbutamide.

3. There was an obvious greater incidence to drug-induced hypoglycemia in the tolbutamide group, as evidenced by the per cent change of fasting blood glucose concentrations from the baseline. Owning to the spontaneous intensification of insulinogenesis and diabetogenic hypoglycemia in many mild diabetics as they do in the U.G.D.P. study, the tolbutamide effect undoubtedly was superimposed upon the individual's own increased elaboration of insulin during the late evening and early morning hours. Administration of the second dose before the evening meal also is pertinent in this context.

4. The basic design of these studies failed to elaborate upon evidences of reactive hypoglycemia either by symptoms, glucose tolerance testing for more than three hours, or concomitant insulin requirements. One would readily assume that a sizable portion of this group did experience significant hypoglycemia, however, on the basis of the fact that on average of 20.4 per cent of all patients studied had a

three-hour blood glucose value of less than 100 mg. per cent.

The tendency in drug-induced hypoglycemia—with oral medication and insulin—is further indicated by the fact that from 37.4 to 49 per cent achieved "good" levels of control—i.e., "70 per cent or more of all the patients showed GTT fasting blood glucose values less than 110 mg. per cent." In a real sense, therefore, the two insulin-treated groups cannot be considered as proper controls in terms of evaluating vascular complications.

In essence, this study underscores the continuing crudeness of our ability to define "control of diabetes." I continually find myself reminding diabetologists that *diabetes mellitus* is NOT a disease of the circulating blood glucose concentration. Rather, it is a disorder of energy metabolism on the tissue level. Accordingly, until we have some practical method for determining the latter, the clinician best advised to define "control of diabetes" to what I have referred to as "enlightened clinical control..." It is my belief, based upon several decades of observation, consultation, and research, that some of the most serious diabetic complications—especially retinopathy and neuropathy—are in small measure attributable to insistence upon "strict control" and the vigorous use of drugs to achieve this goal at the cost of severe tissue glycopenia...

Perhaps the most significant aspect of the U.G.D.P. study is the attention it has focused upon the many deficiencies in our understanding of the pathogenesis and rational treatment of diabetes mellitus. H. J. Ronzans, M.D., West Palm Beach, Fla.

'Dirty Wounds'

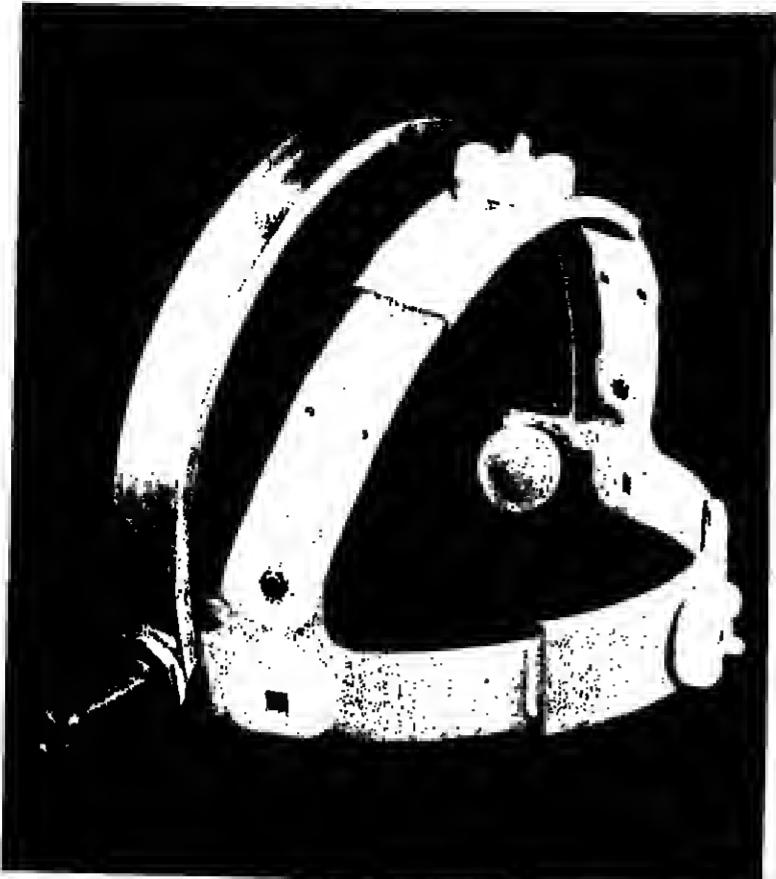
Editor, MEDICAL TRIBUNE:
The action of the larvae of the common housefly (*Musca domestica*) in cleaning up dirty wounds was noted during the first World War.

It was investigated in Britain where the housefly was bred under sterile conditions and the larvae deliberately sown onto dirty wounds. The maggots performed a perfect debridement in that only necrotic and nonviable tissue was removed.

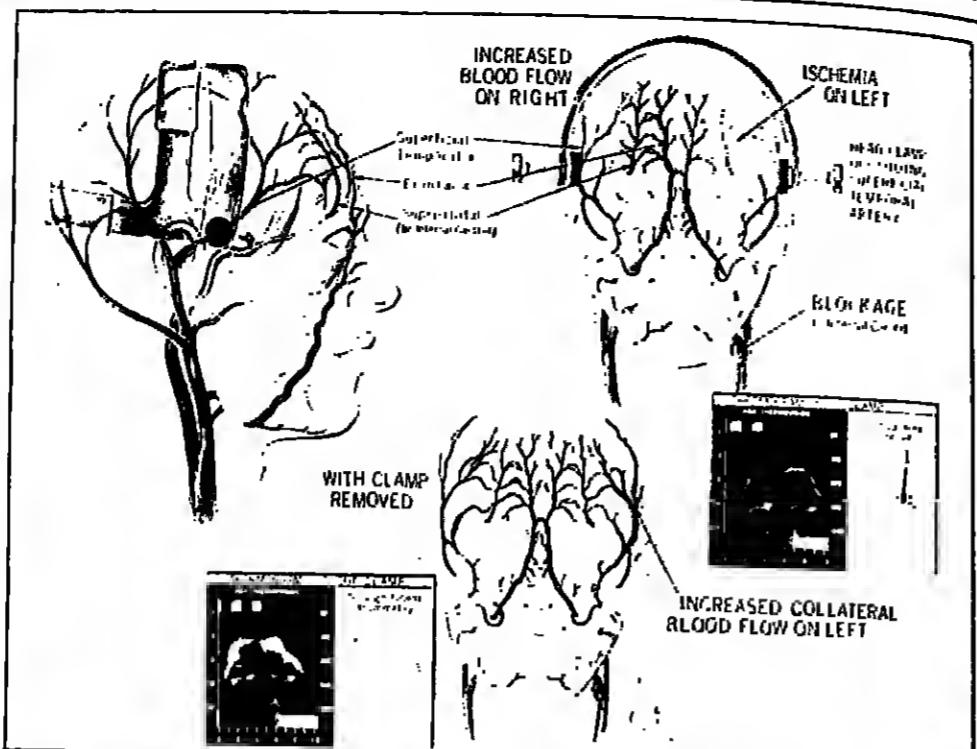
Further investigation showed that the maggots excrete large quantities of urea. Dressing moistened in a saturated aqueous solution of urea were substituted and similar results obtained without the accidentally presence of the larvae.

Dr. Sturto, M.D., D.T.M., Director, Southern Branch Provincial Laboratory of Public Health, Calgary, Alta.

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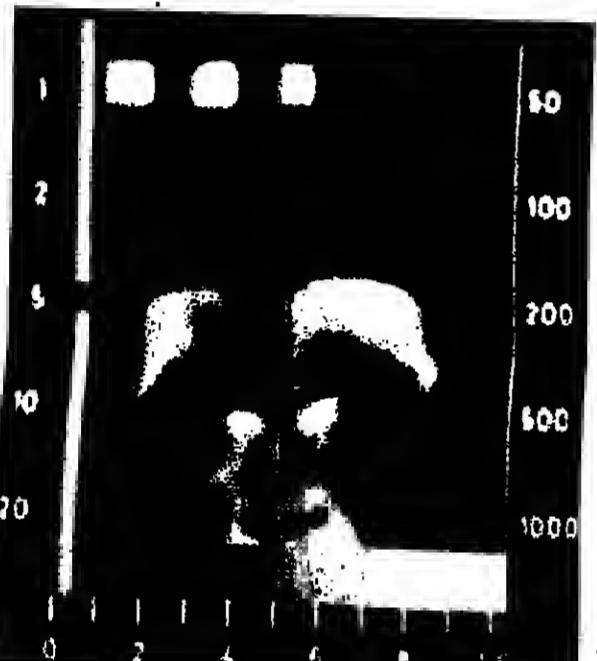
Prototype head clamp has adjustable rubber foot pads to promote bilateral occlusion of the superficial temporal arteries. Investigators took thermograms at least five minutes after application of this device.



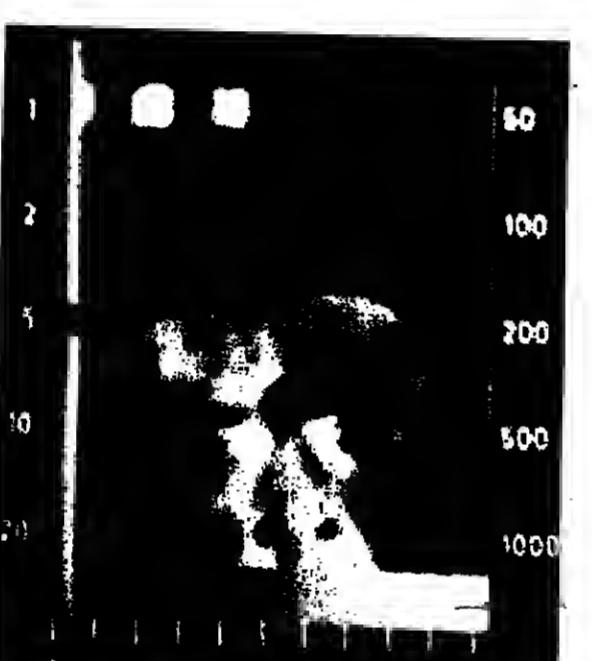
Composite drawing outlines mechanism by which superficial temporary artery clamping reveals hidden abnormalities. In thermogram at right, researchers injected hot water into a cadaver via catheter inserted into left internal carotid artery. Thermograms with forehead temperature asymmetries of 0.7° C. or more were considered abnormal in this study.



Sex-specific differences are major determinants—perhaps greater than environmental factors—of monkey's play behavior, say Wisconsin RPRC investigators. Males, left, seem to prefer contact games; females play independently. Scientists, below, at New England RPRC, perform vectorcardiogram. They cooperate with Boston U. Medical School and MGH Medical Center researchers to study problems of human cardiac disease.



Patient with a left internal carotid stenosis. Thermogram was taken with standard method and does not show significant forehead temperature asymmetry.



Head clamp was utilized in this thermogram of the same patient. Cool area over left eye indicates decreased blood flow in the left internal carotid artery.

Greater Thermogr. Sensitivity Sought Four Basic Faciatterns Discerned In Carotid Occlusive Disease Cases

ASIC PATTERNS of abnormal facial thermograms studied by Dr. Terrance Capistrant, of St. Paul-Ramsey Hospital, St. Paul, to improve sensitivity of this method for predicting carotid occlusive disease. In the project were 315 patients and 200 controls. Of these 35, 31 were detected thermographically. One abnormal thermogram obtained from the latter subjects, by combined head clamp technique, was examined for characteristic features.

Four basic facial patterns were discerned: (1) medial forehead cooling, (2) a horizontal area of the supraorbital ridge that resembled an asymmetric thickening of the skin, (3) cooling of one side of the forehead, and (4) asymmetric cooling in a region supplied by the canthal and palpebral branches of the internal carotid. The latter showed up more often in combination with other findings than alone.

Though some thermograms contained combinations of these, they never comprised more than two types.

The most frequent pattern seen in the head clamp thermograms was category 3, followed by 1. This contrasts with previous findings with standard thermograms, where 1 appeared often and 2 was also very common.



DR. CAPISTRANT

Genetics Counseling Seen as Part Of Trend to Preventive Medicine

GENETICS COUNSELING fits into the present trend towards preventive medicine, stresses Dr. George Solish, Associate Professor of Obstetrics and Gynecology and director of the newly established genetics counseling clinic at Downstate Medical Center, Brooklyn.

A feature of this clinic is close cooperation with other Downstate specialists—such as hematologists and surgeons—made necessary by the great variety of defects (some 1,400 of which are presently known, according to the director) found to have a genetic basis. The unit also serves to monitor congenital malformations due to pollution or drugs taken during pregnancy, says Dr. Solish.

Research and teaching are other functions of the clinic. It is supported by a grant from the Birth Defects Institute of the New York State Department of Health.



Proportional body measurements of 16-year-old with stunted growth are taken by Dr. Solish at the outpatient clinic. In addition, he drew blood samples, took x-rays, and scraped tissue cells from patient's mouth for the purpose of cytogenetic studies.



Dr. Solish's efforts to help the patient.

dermatoglyphic print from mother of 16-year-old boy. The print is being examined by a doctor in a clinical setting.



Union-made. Twice daily, physicians at the University-McCook Hospital in Hartford and the U.S. Veterans Administration Hospital in Newington gather for joint conferences on patients by way of a newly implemented two-way closed-circuit TV system, the first in Connecticut. They exchange questions and comments; charts and graphs can also be transmitted. This hook-up anticipates a network that will ultimately link all of the state's hospitals with the University of Connecticut medical and dental schools and the Yale University School of Medicine.

Primate Research Applied as a Key To Human Health

OVER 100 SCIENTISTS at the seven labs of NIH's Primate Research Centers Program collaborate with visiting investigators in finding solutions for human health problems. Under the institute's Division of Research Resources, these units are situated throughout the U.S. Their locations and school affiliations are Oregon Regional Primate Research Center, Beaverton, University of Oregon; Washington RPRC, Seattle, University of Washington; New England RPRC, Southborough, Mass., Harvard University; Yerkes RPRC, Atlanta, Ga., Emory University; Delta RPRC, Covington, La., Tulane University; National Center for Primate Biology, Davis, University of California; and Wisconsin RPRC, Madison, University of Wisconsin.

Central nervous system, cardiovascular, and neonatal diseases, mental retardation, and reproductive biology are under study.





One to two hours before surgery, 10 mg Injectable Valium (diazepam) I.M.



surroundings and disturbing procedures. Perhaps best of all, Injectable

can promptly calm, lessening anxiety and tension associated with strange

Premedication for surgery

Injectable Valium (diazepam) is a useful premedicant for reducing undue anxiety. Recall of preoperative procedures is markedly diminished. When given in conjunction with narcotics, a reduction of narcotic dosage should be considered. (See summary of prescribing information.) Injectable Valium should not be mixed with other drugs, solutions, or fluids. The new 10-mg disposable syringe can help you observe this precaution at the same time it helps assure aseptic handling. Injectable Valium seldom significantly alters vital signs. Nevertheless, there have been infrequent reports of hypotension and rare reports of apnea and cardiac arrest, usually following I.V. administration. Resuscitative facilities should be available.

To relieve excessive preoperative anxiety, remember Injectable Valium (5 mg/ml) 2-ml ampul, 10-ml vial, 2-ml disposable syringe.

Valium (diazepam) markedly diminishes recall of the preoperative procedure.

Before prescribing, please consult complete product information, a summary of which follows:

Indications: Tension and anxiety states; somatic complaints which are concomitants of emotional factors; psychoaffective states manifested by tension, anxiety, apprehension, fatigue, depressive symptoms or agitation; symptomatic relief of acute agitation, tremor, delirium, trismus and hallucinosis due to acute alcohol withdrawal; adjunctively in: relief of skeletal muscle spasm due to reflex spasm to local pathology; spasticity caused by upper motor neuron disorders; athetosis; stiff-man syndrome; tetanus; status epilepticus and severe recurrent seizures; anxiety

prior to gastroscopy, esophagoscopy, and surgical procedures; cardioversion (I.V.).

Contraindications: In infants; in patients with known hypersensitivity to the drug; in acute narrow angle glaucoma; may be used in patients with open angle glaucoma receiving appropriate therapy.

Warnings: Inject I.V. slowly, directly into vein; take at least one minute for each 5 mg (1 ml) given. Do not mix or dilute with other solutions or drugs. Do not add to I.V. fluids. Rare reports of apnea or cardiac arrest noted, usually following I.V. administration, especially in elderly or very ill and those with limited pulmonary reserve; duration is brief; resuscitative facilities should be

available. Not recommended as sole treatment for psychotic or severely depressed patients. Should not be administered to patients in shock, coma, acute alcoholic intoxication with depression of vital signs. Caution against hazardous occupations requiring complete mental alertness. Advise against simultaneous ingestion of alcohol and other CNS depressants. Withdrawal symptoms (similar to those with barbiturates and alcohol) have occurred following abrupt discontinuance (convulsions, tremor, abdominal and muscle cramps, vomiting and sweating). Keep addiction-prone individuals under careful surveillance because of their predisposition to habituation and dependence. In pregnancy,

lactation or women of childbearing age, weigh potential benefit against possible hazard to mother and child.

Precautions: If combined with other psychotropics or anticonvulsants, carefully consider individual pharmacologic effects—particularly with known compounds which may potentiate action of Valium, such as phenothiazines, narcotics, barbiturates, MAO inhibitors and other antidepressants. Usual precautions indicated in patients severely depressed, or with latent depression, or with suicidal tendencies. Observe usual precautions in impaired renal or hepatic function. Not recommended for bronchoscopy, laryngoscopy, obstetrical use, or in diagnostic procedures other than

gastroscopy and esophagoscopy. Laryngospasm and increased cough reflex are possible during gastroscopy; necessary countermeasures should be available. Hypotension or muscular weakness possible, particularly when used with narcotics, barbiturates or alcohol. Side effect with narcotics may be additive, appropriate reduction in narcotic dosage is possible. Use lower doses (2 to 5 mg) for elderly and debilitated. Safety and efficacy in children under 12 not established.

Side Effects: Drowsiness, fatigue, ataxia, confusion, depression, constipation, dysarthria, diplopia, headache, hypoactivity, hiccup, hypotension, incontinence, jaudice, nausea, changes

in libido, changes in salivation, phlebitis at injection sites, urinary retention, skin rash, syncope, blurred speech, urticaria, tremor, vertigo, blurred vision. Paradoxical reactions such as acute hyperexcited states, anxiety, hallucinations, increased muscle spasticity, insomnia, rage, sleep disturbances and stimulation have been reported; should these occur, use of the drug should be discontinued. Isolated reports of neutropenia, jaundice; periodic blood counts and liver function tests advisable during long-term therapy. Minor EEG changes, usually low-voltage fast activity, of no known significance.



Roche Laboratories
Division of Hoffmann-La Roche Inc.
Nutley, N.J. 07110

Injectable Valium® (diazepam)

benefits every step of the way.

Small Outbreaks Of Salmonellosis Reported in U.S.

Medical Tribune Report

ATLANTA, Ga.—Two outbreaks of salmonellosis—one in Maine and the other in California—were reported here by the Center for Disease Control.

Thirty-three persons attending a christening in Kittery Point, Me., were served lunch on August 7, and within two days 17 of those who ate the meal became ill with gastroenteritis. Cultures of stool specimens from 14 were positive for Salmonella Thompson.

Sixteen persons who ate chicken salad became ill, while only one of 12 who did not eat this food was similarly affected.

Ingredients of the chicken salad included home-grown lettuce and celery, mayonnaise, and three chickens, all purchased at a local supermarket. There were no ingredients or food samples left for laboratory analysis. Environmental cultures were all negative. There was no evident error in food handling or medical history to suggest recent salmonellosis in the food handler.

Fifty per cent of all nonhuman isolates of Salmonella Thompson reported to the CDC in 1970 were obtained from chickens, it was noted. Boiling chicken for one hour should be sufficient to kill salmonellosis. In this case, recontamination presumably occurred after it was boiled, though no specific handling error was documented.

The other outbreak was due to Salmonella Berta and occurred in June in Red Bluff, Calif. Approximately 200 ill persons were identified. Fifteen were hospitalized, and two elderly persons died.

Epidemiologic investigation revealed the vehicle of infection to be custard-filled pastries, particularly maple bars, processed and sold at a single bakery. The contaminated ingredient was unpasteurized,

Asthmatic's Breath Measured



At University Hospitals of Cleveland, Dr. Howard Schwertz, allergist-immunologist, demonstrates breathing measurement device to asthmatic patient. He is investigating effector mechanisms in immune responses.

The most often specified tetracycline* is now one of the least expensive.

Check the price to your patients with your local pharmacist. (You may be surprised)

Achromycin V

Tetracycline HCl—250 mg. capsules, of course

LEADER LABORATORIES A Division of American Cyanamid Company, Pearl River, New York

Exercise to Bar Coronary Ills Held to Need Stress Testing

Continued from page 1

changes his life style, takes reasonable time for rest and relaxation away from his main occupations, finds other outlets that are enjoyable, makes whatever adjustments are necessary and possible in his home situation, one may hope to effect some improvement."

The very least that should be required in the way of stress examination is a Master two-step exercise test and preferably a double Master, Dr. Phibbs said. A monitored multistage exercise study is preferable, he added, and where facilities are available this should always be carried out as a prerequisite for participation in an exercise program, he added.

Respiratory maintenance is the most important part of therapy, he said. Artificial respiration should be given when required, including tracheotomy and positive-pressure ventilation if respiratory failure is not later relieved by antitoxin.

Atropine sulfate should be given intravenously as soon as cyanosis is overcome, but not until then. Intravenous pralidoxime chloride is a specific antitoxin and sometimes gives dramatic results. Finally, other symptomatic treatment can be given and skin, hair, and clothes decontaminated, the toxicologist said.

Respiratory Snags Seldom Tie To Major Black Lung Signs

From W. Va. University

► Coal workers' pneumoconiosis is associated with minor impairments in respiratory function, but by themselves these respiratory deficits are seldom associated with significant symptoms or respiratory disability, the meeting was told by Dr. W. Keith, C. Morgan, N. LeRoy Lips, and Anthony Sinton, of the West Virginia University School of Medicine.

However, the addition of those above maladies to those caused by another mild, unrelated pulmonary disease, such as chronic bronchitis, may result in the patient's developing symptoms that he would not have were only one disease present, they said.

"Exercise in this group of patients improves functional capability in a very clear-cut and striking manner."

Organophosphorus Poisoning May Be on the Increase

From Wenatchee, Wash.

► The medical profession should be prepared to diagnose and treat increased poisoning from organophosphorus pesticides, according to Dr. Griffith E. Quinby, a toxicologist from Wenatchee, Wash.

The restriction of chlorinated hydrocarbons is leading to greater reliance upon the much more toxic organophosphorus compounds, and this will probably lead to an increase in death and morbidity from these chemicals, he explained.

Toxicity of Methotrexate To Liver Is Held Doubtful

Continued from page 1
however, did not differ significantly from untreated psoriasis.

"Results of the study do not suggest abandoning methotrexate as a treatment of severe psoriasis," Dr. Zacharias emphasized, "but may stimulate a wider use of liver biopsy in the control."

The patient denied any known contact with rodents, although chipmunks, tree squirrels, and wood rats are abundant around the cabin area. No die-off of any of these animals was apparent to either the boy or any of his family members. He was unaware of any flea bites and was not certain of the cause of the toe lesion.

The other 10 children and two adults who were living at the cabin have remained well.

Dr. Zacharias remarked that there is no doubt that, in terms of treatment, methotrexate is far less toxic to the liver than continuous treatment to low dosage. His doses were 25 to 50 mg. once weekly, administered intramuscularly.

All of Dr. Zacharias' patients receiving methotrexate displayed an increase in serum glutamic pyruvate transaminase, demonstrating, he commented, that this rise is an almost normal finding during methotrexate treatment.

Chess Solution
White wins by: 1 K-N7, P-OR4; 2 K-R6, P-QR5; 3 K-R5, P-QR6; 4 K-R4, P-QR7; 5 K-N3, threatening 6 N-K6 mate. So Black must play 5...P-K5; 6 KxP, KxP; and eventually all his other pawns will also be captured.

Exercise to Bar Coronary Ills Held to Need Stress Testing

In patients with occupations exposing to organophosphates who present with signs of post-sympathetic overstimulation, a history of overexposure should be obtained but, if necessary, treatment of a supportive nature should begin before the history is obtained, Dr. Quinby recommended. Treatment should never be delayed until laboratory confirmation is obtained, he warned.

Respiratory maintenance is the most important part of therapy, he said. Artificial respiration should be given when required, including tracheotomy and positive-pressure ventilation if respiratory failure is not later relieved by antitoxin.

Atropine sulfate should be given intravenously as soon as cyanosis is overcome, but not until then. Intravenous pralidoxime chloride is a specific antitoxin and sometimes gives dramatic results. Finally, other symptomatic treatment can be given and skin, hair, and clothes decontaminated, the toxicologist said.

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has a system that wins

She has a system that wins. Thanksgiving dinner for eleven. And she handles everything beautifully, wins lots of compliments.

She has another system for her hypertension. And that also works beautifully. It includes Ser-AP-Es.

More than just another antihypertensive, Ser-AP-Es can be a whole medication plan for living with hypertension.

A "recipe" for comfort? Correct. Because Ser-AP-Es controls blood pressure effectively; dosage of each compo-

nent is lower than if prescribed alone, usually minimizing side effects. However, side effects may occur (see brief prescribing information).

Designed with the kidney in mind?

Hydralazine maintains or increases renal blood flow.

And the brain too?

Hydralazine also relaxes cerebral vascular tone. And reserpine has beneficial calming action.

Can she serve herself some "goodies"?

Well, hydrochlorothiazide does eliminate excess salt and water. That may mean less rigid

Ser-AP-Es®

reserpine 0.1 mg
hydralazine hydrochloride 25 mg
hydrochlorothiazide 15 mg

a system for living with hypertension

INDICATIONS: All cases of hypertension except the mildest and the most severe.

CONTRAINDICATIONS: Reserpine: Known hypersensitivity; mental depression, especially with suicidal tendencies; active peptic ulcer; lactation.

Hydralazine: Hypersensitivity to coronary artery disease; severe and/or uncontrolled hypertension; heart disease.

Hydrochlorothiazide: Anuria; progressive renal or hepatic disease; allergy to thiazides or other sulfonamide-derived drugs.

WARNINGS: Reserpine: Withdraw reserpine 2 weeks before surgery, if possible. For emergency surgical procedures, give vagal blocking agents parenterally to prevent or reverse hypotension and/or bradycardia.

Hydralazine: Hypersensitivity to coronary artery disease; severe and/or uncontrolled hypertension; heart disease.

Hydrochlorothiazide: Anuria; progressive renal or hepatic disease; allergy to thiazides or other sulfonamide-derived drugs.

The most often specified tetracycline* is now one of the least expensive.

Check the price to your patients with your local pharmacist. (You may be surprised)

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The ultimate test:

the
recovery
room



Confirms the value of Talwin in the hospital and in private practice

analgesic efficacy comparable to meperidine and morphine with a minimum of significant adverse effects

In postoperative patients: less circulatory depression with Talwin

In a double-blind study of 342 postop patients, circulatory depression occurred in 13.2% of 174 patients receiving morphine, as compared with 5.4% of 168 patients receiving Talwin.¹

...and in other patients: less respiratory and circulatory depression with Talwin

In a double-blind study of 3 narcotic analgesics and Talwin in 118* patients with suspected acute myocardial infarction, Talwin caused a significantly lower incidence in the fall of systolic blood pressure than the 3 narcotics when the initial pressure was 120 mm. Hg or higher. Unlike the narcotics, Talwin caused a statistically significant rise in the systolic blood pressure of patients with initial pressures of less than 120.²

In a study of a series of patients given Talwin or meperidine while anesthetized for surgery, the investigators concluded: "it would therefore appear that pentazocine is a much safer drug in respect of respiratory depression than pethidine [mepidine], particularly when repeated injections are to be given, e.g. postoperatively or in obstetric practice."³

...and less of the other adverse effects associated with narcotic analgesics

Compared to morphine, Talwin is relatively free from urinary retention and constipation.

Is less likely to cause nausea, vomiting and diaphoresis than mepidine.

Available in 3 dosage strengths—all within the range of recommended dosage. Talwin is available in 30 mg., 45 mg., and 60 mg. strengths to provide analgesia specific to patients' needs throughout the course of therapy. Most studies indicate that the higher dosage strengths produce little, if any, increase in the incidence of adverse reactions.

References: 1. Wallace, George: *Int. Surg.* 53:185, Feb. 1970.
2. Scott, M. E. and Orr, Rosemary: *Lancet* 1:1065, May 31, 1969.
3. Davis, L., et al.: *Brit. J. Anesth.* 42:113, Feb. 1970.

*Other drugs studied: dimorphone and meperidine.

Injectable
Talwin®
brand of
pentazocine
(as lactate)
bulwark against
moderate to severe pain



Injectable Talwin® brand of pentazocine (as lactate) Analgesic for parenteral use

- Tolerance to analgesic efficacy has not been observed
- not subject to narcotic controls

Contraindications: Talwin should not be administered to patients who are hypersensitive to it.

Warnings: Head Injury and Increased Intracranial Pressure. The respiratory depressant effects of Talwin and its potential for elevating cerebrospinal fluid pressure may be markedly exaggerated in the presence of head injury, other intracranial lesions, or a preexisting increase in intracranial pressure. Furthermore, Talwin can produce effects which may obscure the clinical course of patients with head injuries. In such patients, Talwin must be used with extreme caution and only at its use is deemed essential.

Use in Pregnancy. Safety in pregnant women during pregnancy (other than labor) has not been established. Animal reproduction studies have not demonstrated teratogenic or embryotoxic effects. However, Talwin should be administered to pregnant patients (other than labor) only when, in the judgment of the physician, the potential benefits outweigh the possible hazards. Patients receiving Talwin during labor have experienced no adverse effects other than those that occur with commonly used analgesics. Talwin should be used with caution in women delivering premature infants.

Drug Dependence. Special care should be exercised in prescribing pentazocine for emotionally unstable patients and for those with a history of drug misuse. Such patients should be closely supervised when long-term therapy is contemplated. There have been instances of psychological and physical dependence on Talwin in patients with such a history and, rarely, in patients without such a history. Abrupt discontinuance following the extended use of parenteral Talwin has resulted in symptoms such as abdominal cramps, elevated temperature, rhinorrhea, restlessness, anxiety, and lacrimation. Even when these occurred, rhinorrhea, restlessness, anxiety, and lacrimation, even when these occurred, discontinuance of Talwin with gradual withdrawal has ameliorated the patient's symptoms. Substituting meperidine or other narcotics for Talwin in the treatment of the Talwin abstinence syndrome should be avoided.

In prescribing parenteral Talwin for chronic use, particularly if the drug is to be self-administered, the physician should take precautions to avoid increases in dose and frequency of injection by the patient and to prevent the use of the drug in anticipation of pain rather than for the relief of pain.

Use in Chronic Pain. Because of clinical experience in children under twelve years of age is limited, the use of Talwin in this age group is not recommended.

Ambulatory Patients. Since sedation, dizziness, and occasional euphoria have been noted, ambulatory patients should be warned not to operate machinery, drive cars, or unnecessarily expose themselves to hazards.

Presumptions: Certain Respiratory Conditions. The possibility that Talwin may cause respiratory depression should be considered in treatment of patients with bronchial asthma. Talwin should be administered only with caution and in low dosage to patients with respiratory depression (e.g., from other medication, uremia, or severe infection), obstructive respiratory conditions, or cyanosis. **Impaired Renal Function.** Although Talwin is extensively metabolized, it is believed that Talwin causes or increases renal or hepatic impairment. The drug should be administered with caution to patients with such impairment. Extensive liver disease appears to predispose to greater side effects (e.g., marked apprehension, anxiety, dizziness, sleepiness) from the usual clinical dose, and may be the result of decreased metabolism of the drug by the liver.

Myocardial Infarction. As with all drugs, Talwin should be used with caution in patients with myocardial infarction who have nausea or vomiting.

Birth Surgery. The safety of Talwin in pregnant patients is unknown. The safety of Talwin in children is unknown. The drug should be used with caution in patients about to undergo surgery of the biliary tract.

Patients Receiving Narcotics. Talwin is a mild narcotic analgesic. Some patients previously receiving narcotics have experienced mild withdrawal symptoms after receiving Talwin.

CNS Effect. Caution should be used when Talwin is administered to patients prone to seizures. Seizures have occurred in a few such patients in association with the use of Talwin, although no cause and effect relationship has been established.

Adverse Reactions: The most commonly occurring reactions are: nausea, dizziness, or light-headedness, vomiting, euphoria. Infrequently occurring reactions are—respiratory: respiratory depression, dyspnea, transient spasm in a small number of newborn infants whose mothers received Talwin during labor; cardiovascular: circulatory depression, shock, hypertension; CNS effects: sedation, alteration of mood (nervousness, apprehension, depression, feeling "feeling"), dreams, gastritis/nausea; constipation, dry mouth; dermatologic (including focal): diaphoresis, sting on injection, flushed skin including phlebitis, dermatitis including pruritis; other: urinary retention, headache, peripheral edema, alterations in rate of strength of uterine contractions during labor.

Rarely reported reactions include—neuromuscular and psychiatric: muscle tremor, incontinence, disorientation, hallucinations; gastrointestinal: taste alteration, diarrhea and cramps; ophthalmic: blurred vision, nystagmus, diplopia, miosis; other: tachycardia, nodules and ulceration at injection site, weakness or lethargy, chills, moderate transient eosinophilia, allergic reactions including edema of the face.

See Acute CNS Manifestations and Drug Dependence under WARNINGS.

Dosage and Administration: Adults (Excluding Patients in Labor). The recommended parenteral dose is 30 mg. by intramuscular, subcutaneous or intravenous route. This may be repeated every 3 to 4 hours. Doses in excess of 30 mg. intravenously or 60 mg. intramuscularly or subcutaneously are not recommended. Total daily dosage should not exceed 360 mg. As with most parenteral drugs, when frequent daily injections are needed over a prolonged period, intramuscular administration is preferable to subcutaneous. In addition, constant rotation of injection sites (e.g., the upper outer quadrant of the buttocks, mid-lateral aspects of the thighs, and the deltoid area) is recommended.

Pain in Labor. A single, 30 mg. parenteral dose of Talwin may be commonly administered. An intravenous 20 mg. dose has given adequate pain relief to some patients in labor when contractions become regular, and this dose may be given two or three times at two- to three-hour intervals, as needed.

Children Under 12 Years of Age. Since clinical experience in children under twelve years of age is limited, the use of Talwin in this age group is not recommended.

CAUTION: Talwin should not be mixed in the same syringes with soluble barbiturates because precipitation will occur.

Overdosage, Manifestations: Children in pain with Talwin overdose has been insufficient to determine the limits of the condition.

Treatment: Oxygen, intravenous fluids, vasopressors, and other supportive measures should be employed as indicated. Assisted or controlled ventilation should also be considered. Although neophentyl and levophenyl are not allelostatics for respiratory depression due to overdosage or unusual sensitivity to Talwin, parenteral naloxone (Narcan® available through Endo Laboratories) is a specific and effective antagonist. If naloxone is not available, parenteral administration of the anticholinergic, methylphenidate (Ritalin®), may be of value if respiratory depression is present.

Talwin is not subject to narcotic controls.

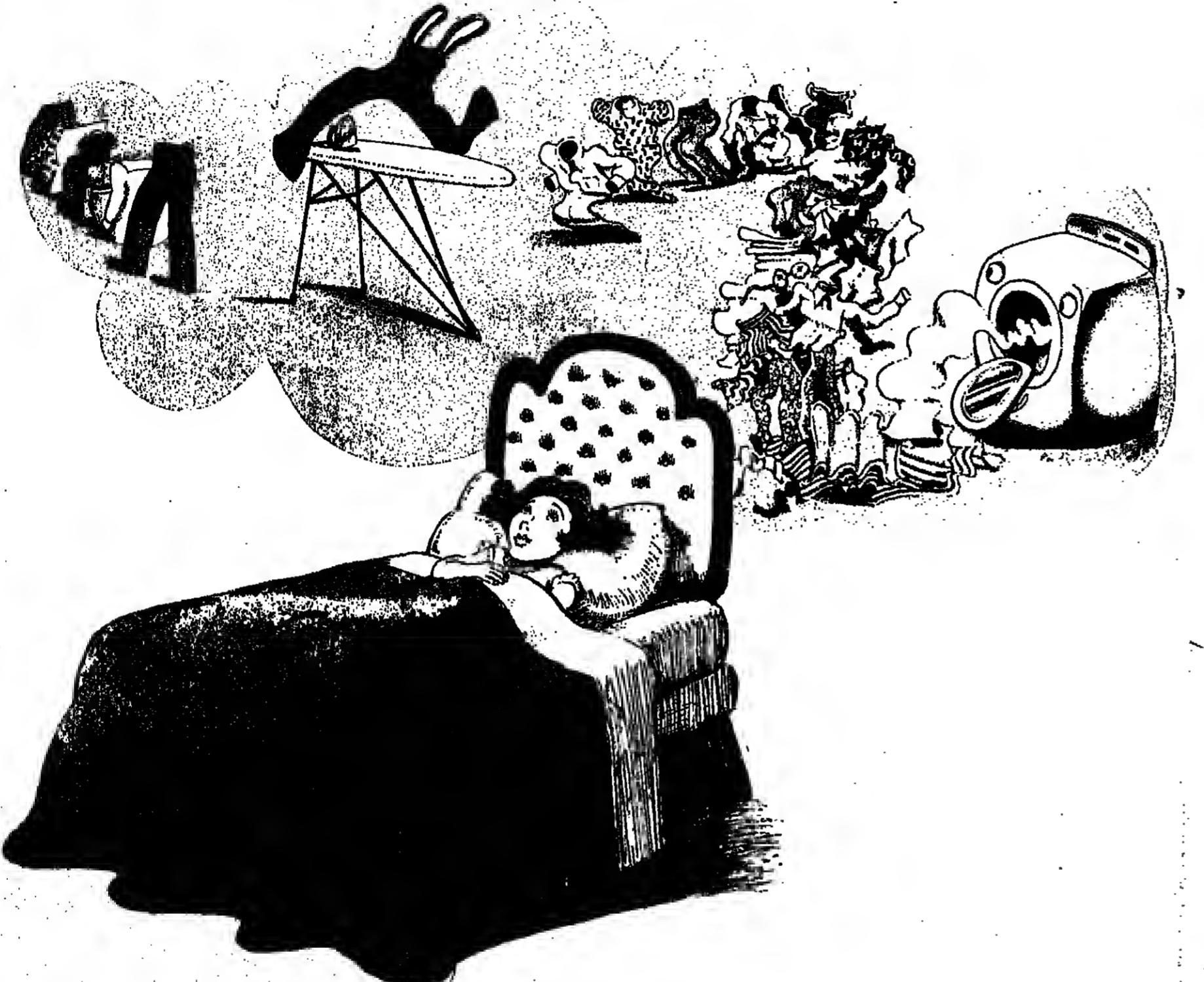
How Supplied: Ampule of 1 ml. (30 mg.), 1/4 ml. (45 mg.), and 2 ml. (60 mg.) each 1 ml. containing Talwin (brand of pentazocine) as lactate equivalent to 30 mg. base, and 2.8 mg. sodium chloride, in water for injection. Boxes of 10, 25, and 100.

Multiple dose vials of 10 ml., each 1 ml. containing Talwin (brand of pentazocine) as lactate equivalent to 30 mg. base, 2 mg. sodium acidulite, 1.8 mg. sodium chloride, and 1 mg. methylparaben as preservative, in water for injection.

The pH of Talwin solutions is adjusted between 4 and 5 with lactic acid and sodium hydroxide. The air in the ampule and vials has been displaced with nitrogen gas.

Winthrop Winthrop Laboratories, New York, N.Y. 10016

Doriden® (glutethimide) and you can count on the rest



At night, she wants to wash her hands of the whole household routine. But there's always another shirt around the corner.

And she finds herself counting them like the proverbial sheep to overcome her insomnia.

It could be so much easier with Doriden. Because Doriden works nice and easy. Usually brings sleep quickly and smoothly. Preeexcitation is rarely a problem.

What's more, Doriden wears off soon, enough so your patient usually wakes naturally refreshed. Morning hangover is also rare.

And Doriden is generally well tolerated by the aged, the chronically ill or hospitalized, those with renal or pulmonary dysfunction—practically anybody with insomnia.

So with Doriden you both rest assured.

INDICATIONS: For night-time, daytime, and preoperative sedation, as well as during first stage of labor.

CONTRAINDICATIONS: Known hypersensitivity to glutethimide.

WARNINGS: Caution patients about possible combined effects with alcohol and other CNS depressants. Do not operate machinery, drive motor vehicle, or engage in activities requiring complete alertness shortly after ingesting drug.

DOSAGE: To avoid over sedation, individualized adjustments during and on cessation of glutethimide therapy.

Physical and Psychological Dependence: Physical and psychological dependence have occurred. Prescribe cautiously for patients known to take excessive quantities of drugs.

Limits: repeated prescriptions without adequate medical supervision. Withdrawal symptoms include nausea, abdominal discomfort, tremors, convulsions, and delirium.

Newborn infants of mothers dependent on glutethimide may also exhibit withdrawal symptoms. In the presence of dependence, dosage should be reduced gradually.

Pregnancy: Use of any drug in pregnancy or lactation requires weighing potential benefits against hazards.

PRECAUTIONS: Total daily dosage above 1 Gm is not recommended for continued administration. In presence of pain, which may counteract the sedative effect of glutethimide, an analgesic should also be prescribed.

ADVERSE REACTIONS: Withdrawal glutethimide if a generalized skin rash occurs. Rash usually clears spontaneously within a few days after withdrawal. Occasional

ally, a purpuric or urticarial rash may occur; exfoliative dermatitis has been reported rarely. With recommended doses, there have been rare reports of nausea, hangover, paradoxical excitation, and blurring of vision. Rarely, acute hypersensitivity reactions, porphyria, and blood dyscrasias (thrombocytopenic purpura, aplastic anemia, leukopenia) have been reported.

DOSAGE: To avoid over sedation, individualized adjustments during and on cessation of glutethimide therapy.

Night-time sedation: 0.25 to 0.5 Gm at bedtime. Repeat dose if necessary, but not less than 4 hours before arising.

Daytime sedation: 0.125 to 0.25 Gm t.i.d. after meals.

Preoperative sedation: 0.5 Gm the night before surgery; 0.5 to 1 Gm t hour before anesthesia. **First stage of labor:** 0.5 Gm at onset of labor. Repeat if necessary.

Supplied: Tablets, 0.5 Gm (white, scored); bottles of 100, 500, 1000 and Script Dispensers of 100. Tablets, 0.25 Gm (white, scored); bottles of 100 and 1000. Tablets, 0.125 Gm (white); bottles of 100. Capsules, 0.5 Gm (blue and white); bottles of 100.

Consult complete literature before prescribing.

CIBA Pharmaceutical Company
Division of CIBA-GEIGY Corporation
Summit, New Jersey 07901

C I B A

Wednesday, September 29, 1971

MEDICAL TRIBUNE

23

The Mail

• "Heart Watch Urged in Vital Hepatitis," says a cover of *Internist Observer*, sent to us by Dr. Edgar M. Allen of Ward, Wash., who wants to know if that's the nonfatal kind.

• The high moral standards of precision machinery were the subject of a story in the *Los Angeles Times* that was forwarded to us by Dr. Albert Fields of that city.

"Dial Manufacturing Co., a precision machine shop, does not have dealing in pornography," it began, and ended by explaining that the company had leased the premises to another company, which had sublet part of it "to others who apparently engaged in activities not compatible with a precision machine shop."

• Child (1). An old friend has turned up in two more places. Dr. Nellie D. Wright of Bristol, Va., found it in *J.A.M.A.*

"All states, starting with the early 1800's passed laws against abortion except when necessary to save the life of the mother, (or in some few cases to protect her health or to save the life of the child.)"

Dr. Howard R. Grove of Bakersfield, Calif., found it in *American Medical News*:

"In Washington, abortion is permitted only to preserve the life of the mother or that of the child."

• Child (2). Dr. Sam A. Nixon of Floresville, Tex., on the other hand, found an odd type of neonate in an instructional ad on VD in *Family Health*:

"To one small city with a population of less than 30,000 one case alone, when traced out, led to the identification and cure of over one hundred and forty-six people including one congenital newborn."

• Readers old enough to remember the era of double-talk should be given a touch of nostalgia by a lucky find in *American Medical News* that was made by Dr. Alan E. Van Scher of Larchmont, N.Y.

"William W. Travis of Alabaster, Ala., prenames 'renders' included hospital pharmacists...."

They're the ones who dispense pharmacies, obviously.

• Drs. Charles E. Jaekla of Defiance, Ohio, and T. Stacy Lloyd, Jr., of Fredericksburg, Va., were not overimpressed with *MEDICAL TRIBUNE* headline that said: "Psychiatric Ills Common in Young Adults Who Tried Suicide."

"Profound observation," wrote Dr. Jaekla; "So what else is new?" asked Dr. Lloyd; "Well, that's what the silly story said," said the chap responsible for the headline.

• "Doo Rickles' wife gave birth to a Cedars of Lebanon Hospital. He is the couple's second child," says the item in the Santa Barbara, Calif., *News-Press* that J. F. Smithson, D.V.M., sent in.

Now there's a cause for sibling rivalry for the firstborn!

• Breakthrough Dr. Mark J. Popp of Brookfield, Wis., contributed the following prize award announcement from the minutes of the annual business meeting of the American College of Obstetricians and Gynecologists. The prize went to an exhibitor entitled: "The Use of Anti-RH Antibiotics for the Prevention of RH Immunization."

On the seventh day of illness the child was admitted to the hospital, still complaining of stomach pain, and, after a blood test and x-rays, underwent surgery, which disclosed a ruptured appendix and peritonitis. The infection was drained, and two months later, the appendix was removed. About a month later, the child was brought to the physician for treatment of a penicillin reaction and a bowel obstruction.

• Dr. Herbert Notkin of Philadelphia writes us:

"There is a Public Health Service department with the following name:

"Institutional Assurance on Investigations Involving Human Subjects, Including Clinical Research and Investigations in Behavioral and Social Sciences."

It's probably known as Freddy to the in-group.

Readers are invited to contribute items of 100 words or less to this column. Contributions should be mailed to MEDICAL TRIBUNE, 110 East 39th St., New York, N.Y. 10022.

Knee Injury Rate Much Higher In Footballers Harmed Before

Medical Tribune Report

New York—A four-year study of more than 61,000 varsity high school football players shows that those with previous serious knee injury are more apt to be seriously reinjured than players with sound knees, it was reported here at a meeting of the Medical Society of the State of New York Committee on the Medical Aspects of Sports.

The study clearly implicates the use of long cleats on the sole of the shoe and finds that elimination of such cleats may reduce the number of knee injuries by half, according to William T. Callahan, Ph.D., director of the study, which was initiated by the New York State Public High School Athletic Association.

"With regard to reducing the seriousness of injuries to the lower extremities, particularly the knee," he said, "the data indicated clearly that two equipment combinations—low shoe, disk heel, ankle wrap and low shoe, short cleat, and no ankle support—produced a statistically significant and lower rate of serious injuries. Conversely, a combination of low shoe, conventional cleats, and ankle tape produced a greater number of serious knee injuries than expected."

"It is recommended, therefore, that high school football players be outfitted with low shoes and some form of disk, or flat heels, or short cleats. It is further recommended that a similar modification—shortening—of the sole cleats on football shoes be made as soon as possible."

The study, conducted by Dr. Callahan

in association with Francis Crowley, Ph.D., Fordham University, and J. Kenneth Hafner, of the New York State Public High School Athletic League, also proposed that players who suffer a serious knee injury be required to take part in a planned program of rehabilitation under the direction of a physician as a precondition to further varsity football competition. Furthermore, such players should be rigorously examined by a physician at the start of each season to determine the degree of rehabilitation of the injured knee and ultimate fitness of the player to participate in competition.

An Evolutionary Process

Discussing the study's findings, Dr. Callahan said that the reduction of serious injuries to the lower extremities is, and will be, an evolutionary rather than a revolutionary process.

"For example," he continued, "if all players were immediately equipped with low shoes and drastically shortened cleats (heel and sole), the total number of serious knee injuries (328) would be reduced by only some 10-20 cases. Furthermore, if all players with a history of previous serious knee injuries were completely rehabilitated and returned to play, the total of serious knee injuries would only be reduced, state-wide, by another 40-50 cases."

"By means of simple arithmetic, it can be seen that some 258 serious knee injuries could still be expected during the course of a varsity high school football season. Elimination of long cleats on the soles of the shoe [and of cleats on the heel] will probably cut that total in half—perhaps to some 130 serious injuries."

Other recommendations:

• That more attention be given to the relative levels of physical maturity of the youngest and oldest boys eligible to compete in varsity play.

• That year-round programs of general

Appointed at Harvard



Dr. Robert T. McCluskey has been appointed S. Burt Wolbach Professor of Pathology by Harvard Medical School. He is also pathologist-in-chief at the Children's Hospital Medical Center.

physical fitness for all high school students, athletes and noncompetitors, be conducted.

• That year-round conditioning programs be directed to the areas of highest vulnerability, such as the knee.

• That extended spring or summer pre-season conditioning programs be instituted for football players.

Dr. Callahan stressed that the results of the study indicate there is a great need for changing the rules of high school football to modify or eliminate potentially dangerous situations and also a need for upgrading the understanding and skills of game officials.

"Certain techniques, such as 'crack back' blocking, piling on, clipping, spearing, 'blind side' blocking, and the like, must be discouraged by coaches and officials," he said. "Field zones in which clipping is legal must be restricted and innovations designed to improve safety—restriction of blocking to above the waist, for example—must be tried and evaluated regardless of whether [or not] they change the 'traditional' nature of the game."

Medicolegal Report

Appendicitis Misdiagnosis Is Not Penalized

Medical Tribune Report

CHICAGO—A physician who misdiagnosed appendicitis as tonsillitis was not negligent, the Supreme Court of Iowa ruled, noting that the symptoms and x-ray findings were consistent with both diseases.

The patient, a five-year-old child, had vomiting, abdominal pain in the lower right side, and a temperature of 101° F.

and was brought to the hospital by her mother. She told the examining physician, who had not seen the child before, that her daughter had appendicitis, but the physician, after examination and the taking of x-rays, made a diagnosis of tonsillitis and sent the child home with medication.

The radiologist, who had reported to the physician his impression from the x-rays that there was localized reflex ileus secondary to appendicitis, testified on the difference between an impression and a diagnosis, and he stated that many disorders in addition to appendicitis could be consistent with reflex ileus, including tonsillitis.

The court said that a physician does not

amine her abdomen without finding sufficient signs for a diagnosis of appendicitis. He had not got a good look at her tonsils, but had noticed that the roof of her mouth was red, and he said that her white blood cell count was 5,000, consistent with tonsillitis.

The radiologist, who had reported to the physician his impression from the x-rays that there was localized reflex ileus secondary to appendicitis, testified on the difference between an impression and a diagnosis, and he stated that many disorders in addition to appendicitis could be consistent with reflex ileus, including tonsillitis.

The court said that a physician does not

insure the correctness of his diagnosis and

that in the present case laymen could not conceive of the complex nature of the diagnostic problem. It held that there was no evidence of incorrect interpretation of the x-rays and that the physician had no duty to advise the parents that the condition seen on the x-rays was also consistent with appendicitis.

It further held that the physician and the radiologist were independent contractors, not hospital employees, and the hospital was not required to inform the patient of the radiologist's impression. The radiologist's reports were made to the physician who was responsible to the parents for the proper handling of the case. (*Shimley v. Surgical Associates*, [186 N.W. 2d 658 (Iowa Sup. Ct., May 5, 1971)].)

MEDICAL MEETING SCHEDULE

Domestic Meetings

Oct. 18-22	American College of Surgeons, Atlantic City, N.J.
Oct. 18-22	Society for Applied Spectroscopy, St. Louis, Mo.
Oct. 19-22	American Society for Microbiology, Atlantic City, N.J.
Oct. 21-23	American Academy of Clinical Toxicology, Philadelphia.
Oct. 21-23	National Hemophilia Foundation, Cleveland.
Oct. 23-25	Nevada State Medical Association, Las Vegas.
Oct. 23-25	American Society of Clinical Pathologists, Boston.
Oct. 23-25	Congress of American Pathologists, Boston.
Oct. 23-26	National Practice Management and Investment Seminar, 45th Annual Meeting, Honolulu.
Oct. 23-27	Eastern Orthopaedic Association, White Sulphur Springs, W. Va.
Oct. 24-26	American College of Chest Physicians, Philadelphia.
Oct. 24-30	American College of Gastroenterology, Atlanta, Ga.
Oct. 29	Society of Teachers of Family Medicine, Washington.
Oct. 30	American Medical Tennis Association, Las Vegas, Nev.
Oct. 31	Academy of Psychosomatic Medicine, Phoenix, Ariz.
Oct. 31	Congress of American Radiologists, Phoenix, Ariz.
Nov. 1	College of Radiologists, Philadelphia.
Oct. 30	Society of Teachers of Family Medicine, Washington.
Oct. 30	American Medical Tennis Association, Las Vegas, Nev.
Oct. 31	Academy of Psychosomatic Medicine, Phoenix, Ariz.
Oct. 31	Congress of American Radiologists, Phoenix, Ariz.
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